

STIC-EIC1600/2900

261002

LCS

From: JON ANGELL [JonE.Angell@uspto.gov]  
Sent: Tuesday, May 20, 2008 1:45 PM  
To: STIC-EIC1600/2900  
Cc: NPL Feedback  
Subject: Database Search Request, Serial Number: 10/696488

Requester: JON ANGELL (P/1635)  
Att. Unit: P/1635  
Employee Number: [REDACTED]  
Office Location: REM 02A05  
Phone Number: (571)272-0756  
Mailbox Number: 2C18

Case serial number: 10/696488  
Class / Subclass(es): 514/44  
Earliest Priority Filing Date: 5/27/1997  
Format preferred for results: Paper  
Attachment: Yes,  
Search Topic Information:

Please search for the structure of formula (I) in claim 1 where A is C(H)(R3)-N(R1)(R2) wherein R1=H, R2=H, R3=H, as indicated in the 4/3/2008 response to the election (attached).  
Special Instructions and Other Comments:

I am available 9:30-5:30 M-F at 2-0756. If possible, I would like to have the search results by June 9.

RECEIVED  
MAY 20 2008  
STIC-EIC1600/2900

5/20/2008

=&gt; fil cap

FILE 'CAPLUS' ENTERED AT 16:49:03 ON 23 MAY 2008

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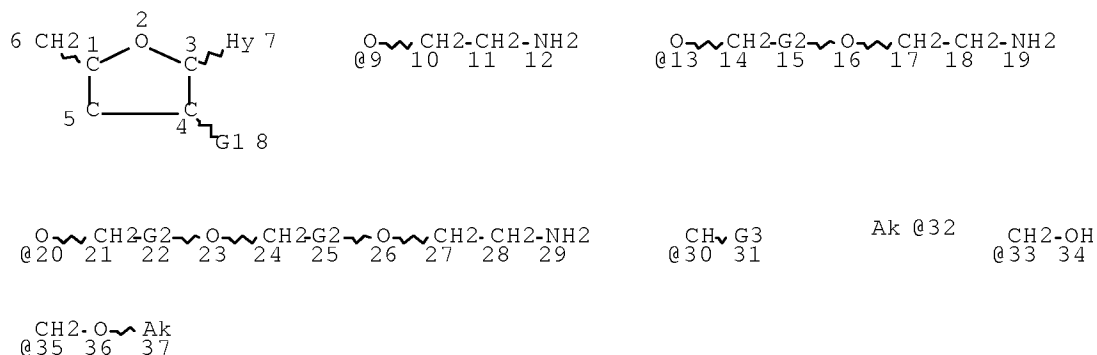
FILE COVERS 1907 - 23 May 2008 VOL 148 ISS 22  
FILE LAST UPDATED: 22 May 2008 (20080522/ED)

Effective October 17, 2005, revised CAS Information Use Policies apply. They are available for your review at:

<http://www.cas.org/legal/infopolicy.html>

=> d que l4

L1 STR



VAR G1=9/13/20

VAR G2=CH2/30

VAR G3=32/33/35

NODE ATTRIBUTES:

CONNECT IS E1 RC AT 32

CONNECT IS E1 RC AT 37

DEFAULT MLEVEL IS ATOM

GGCAT IS LOC SAT AT 32

GGCAT IS LOC SAT AT 37

DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED

NUMBER OF NODES IS 37

STEREO ATTRIBUTES: NONE

L3 4 SEA FILE=REGISTRY SSS FUL L1

L4 9 SEA FILE=CAPLUS ABB=ON PLU=ON L3

=> fil wpix

FILE 'WPIX' ENTERED AT 16:49:13 ON 23 MAY 2008

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FILE LAST UPDATED: 19 MAY 2008 <20080519/UP>  
 MOST RECENT THOMSON SCIENTIFIC UPDATE: 200832 <200832/DW>  
 DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

>>> IPC Reform backfile reclassifications have been loaded to the end of March 2008. No update date (UP) has been created for the reclassified documents, but they can be identified by 20060101/UPIC and 20061231/UPIC, 20070601/UPIC, 20071001/UPIC, 20071130/UPIC and 20080401/UPIC.  
 ECLA reclassifications to April and US national classifications to the end of January 2008 have also been loaded. Update dates 20080401/UPEC and /UPNC have been assigned to these. <<<

FOR A COPY OF THE DERWENT WORLD PATENTS INDEX STN USER GUIDE,  
 PLEASE VISIT:  
[http://www.stn-international.de/training\\_center/patents/stn\\_guide.pdf](http://www.stn-international.de/training_center/patents/stn_guide.pdf)

FOR DETAILS OF THE PATENTS COVERED IN CURRENT UPDATES, SEE  
<http://scientific.thomsonreuters/support/patents/coverage/latestupdates/>

EXPLORE DERWENT WORLD PATENTS INDEX IN STN ANAVIST, VERSION 2.0:  
[http://www.stn-international.com/archive/presentations/DWPIAnaVist2\\_0710.pdf](http://www.stn-international.com/archive/presentations/DWPIAnaVist2_0710.pdf)

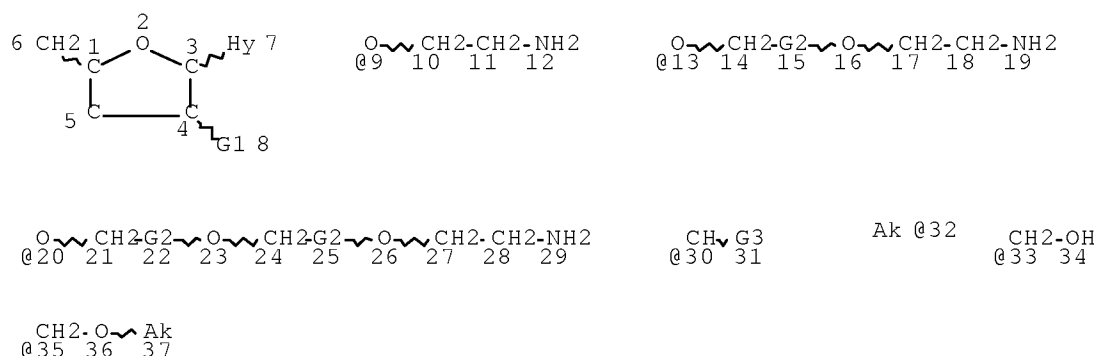
>>> HELP for European Patent Classifications see HELP ECLA, HELP ICO <<<

>>> Updated PDF files in the following links:  
[http://www.stn-international.de/stndatabases/details/ico\\_0803.zip](http://www.stn-international.de/stndatabases/details/ico_0803.zip)  
[http://www.stn-international.de/stndatabases/details/epc\\_0803.zip](http://www.stn-international.de/stndatabases/details/epc_0803.zip)  
 Supplement of all changed ECLA items:  
[http://www.stn-international.de/stndatabases/details/ecla\\_0804s.zip](http://www.stn-international.de/stndatabases/details/ecla_0804s.zip) <<<

>>> Please note that the COPYRIGHT notification has changed <<<

=> d que 17

L1 STR



VAR G1=9/13/20

VAR G2=CH<sub>2</sub>/30

VAR G3=32/33/35

NODE ATTRIBUTES:

CONNECT IS E1 RC AT 32

CONNECT IS E1 RC AT 37

DEFAULT MLEVEL IS ATOM  
 GGCAT IS LOC SAT AT 32  
 GGCAT IS LOC SAT AT 37  
 DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:  
 RING(S) ARE ISOLATED OR EMBEDDED  
 NUMBER OF NODES IS 37

STEREO ATTRIBUTES: NONE

L6 1 SEA FILE=WPIX SSS FUL L1  
 L7 1 SEA FILE=WPIX ABB=ON PLU=ON L6/DCR

=> fil marpat

FILE 'MARPAT' ENTERED AT 16:49:19 ON 23 MAY 2008  
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FILE CONTENT: 1961-PRESENT VOL 148 ISS 20 (20080516/ED)

SOME MARPAT RECORDS ARE DERIVED FROM INPI DATA FOR 1961-1987

MOST RECENT CITATIONS FOR PATENTS FROM MAJOR ISSUING AGENCIES  
 (COVERAGE TO THESE DATES IS NOT COMPLETE):

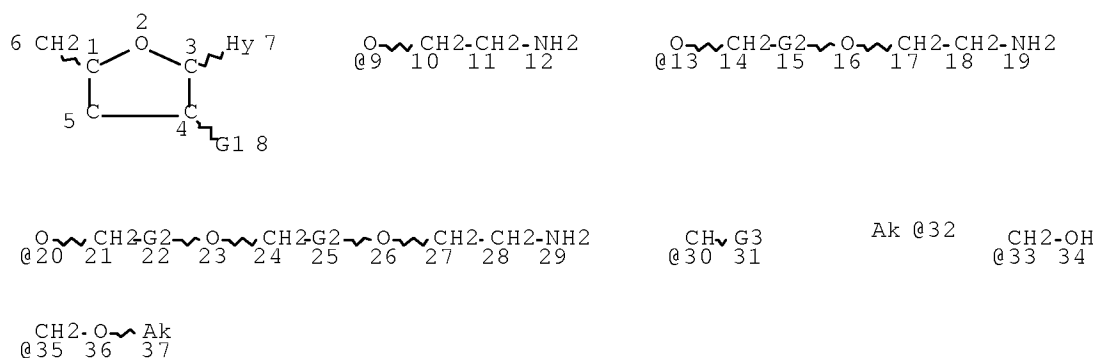
US 20080081929 03 APR 2008  
 DE 102006045603 03 APR 2008  
 EP 1905420 02 APR 2008  
 JP 2008072059 27 MAR 2008  
 WO 2008040000 03 APR 2008  
 GB 2441892 19 MAR 2008  
 FR 2906251 28 MAR 2008  
 RU 2321037 27 MAR 2008  
 CA 2611532 08 MAR 2008

Expanded G-group definition display now available.

Effective December 15th the iteration and answer limits in MARPAT have increased from 100,000 to 200,000 for both on-line and batch searches. For more information on MARPAT search limits, type HELP SLIMITS at an arrow prompt.

=> d que 112

L1 STR



VAR G1=9/13/20  
VAR G2=CH2/30  
VAR G3=32/33/35  
NODE ATTRIBUTES:  
CONNECT IS E1 RC AT 32  
CONNECT IS E1 RC AT 37  
DEFAULT MLEVEL IS ATOM  
GGCAT IS LOC SAT AT 32  
GGCAT IS LOC SAT AT 37  
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:  
RING(S) ARE ISOLATED OR EMBEDDED  
NUMBER OF NODES IS 37

STEREO ATTRIBUTES: NONE  
L12 17 SEA FILE=MARPAT SSS FUL L1

=> dup rem l4 l7 l12  
FILE 'CAPLUS' ENTERED AT 16:49:24 ON 23 MAY 2008  
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FILE 'MARPAT' ENTERED AT 16:49:24 ON 23 MAY 2008  
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COPYRIGHT (C) 2008 American Chemical Society (ACS)  
PROCESSING COMPLETED FOR L4  
PROCESSING COMPLETED FOR L7  
PROCESSING COMPLETED FOR L12  
L13 27 DUP REM L4 L7 L12 (0 DUPLICATES REMOVED)  
ANSWERS '1-9' FROM FILE CAPLUS  
ANSWER '10' FROM FILE WPIX  
ANSWERS '11-27' FROM FILE MARPAT

=> d l13 ibib abs hitstr 1-10;d l13 ibib abs qhit 11-27

L13 ANSWER 1 OF 27 CAPLUS COPYRIGHT 2008 ACS on STN  
ACCESSION NUMBER: 2007:238826 CAPLUS Full-text  
DOCUMENT NUMBER: 148:331943  
TITLE: N,N'-bis-(2-(cyano)ethoxycarbonyl)-2-methyl-2-thio-  
pseudo-urea: a guanylation reagent for synthesis of  
2'-O-[2-(guanidinium)ethyl]-modified oligonucleotides  
AUTHOR(S): Prakash, Thazha P.; Puschl, Ask; Manoharan, Muthiah  
CORPORATE SOURCE: Department of Medicinal Chemistry, Isis  
Pharmaceuticals Inc., Carlsbad, CA, USA  
SOURCE: Nucleosides, Nucleotides & Nucleic Acids (2007),  
26(2), 149-159  
CODEN: NNNAFY; ISSN: 1525-7770  
PUBLISHER: Taylor & Francis, Inc.  
DOCUMENT TYPE: Journal  
LANGUAGE: English

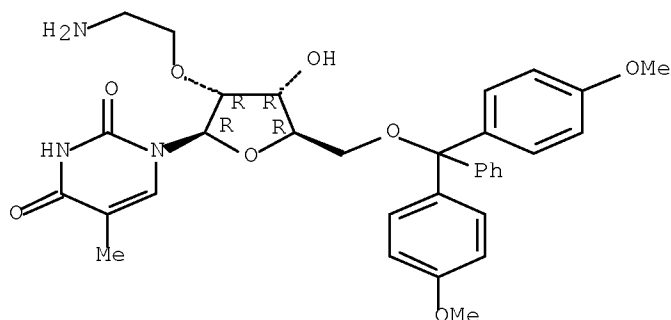
AB A guanylyating reagent, N,N'-bis-(2-(cyano)ethoxycarbonyl)-2-thio-pseudo- urea, was synthesized and used for synthesis of 2'-O-[2- (guanidinium)ethyl] (2'-O-GE) modified oligonucleotides via cyclization reaction. A convenient deprotection method for the 2'-O-GE oligonucleotides was developed.

IT 242147-79-1  
 RL: RCT (Reactant); RACT (Reactant or reagent)  
 (N,N'-bis-(2-(cyano)ethoxycarbonyl)-2-methyl-2-thio-pseudo-urea as guanylyating reagent for synthesis of 2'-O-[2-(guanidinium)ethyl]-modified oligonucleotides)

RN 242147-79-1 CAPLUS

CN Uridine, 2'-O-(2-aminoethyl)-5'-O-[bis(4-methoxyphenyl)phenylmethyl]-5-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 19 THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 2 OF 27 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2006:67953 CAPLUS [Full-text](#)

DOCUMENT NUMBER: 144:327203

TITLE: Fluorescence of covalently attached pyrene as a general RNA folding probe

AUTHOR(S): Smalley, Mary K.; Silverman, Scott K.

CORPORATE SOURCE: Department of Chemistry, University of Illinois at Urbana-Champaign, Urbana, IL, 61801, USA

SOURCE: Nucleic Acids Research (2006), 34(1), 152-166

CODEN: NARHAD; ISSN: 0305-1048

PUBLISHER: Oxford University Press

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Fluorescence techniques are commonly and powerfully applied to monitor biomol. folding. In a limited fashion, the fluorescence emission intensity of covalently attached pyrene has been used as a reporter of RNA conformational changes. Here, the authors pursue two goals: the authors examine the relation between tether identity and fluorescence response, and the authors determine the general utility of pyrene fluorescence to monitor RNA folding. The P4-P6 domain of the Tetrahymena group I intron RNA was systematically modified at multiple nucleotide positions with pyrene derivs. that provide a range of tether lengths and compns. between the RNA and chromophore. Certain tethers typically lead to a superior fluorescence signal upon RNA folding, as demonstrated by equilibrium titrns. with Mg2+. In addition, useful fluorescence responses were obtained with pyrene placed at several nucleotide positions dispersed throughout P4-P6. This suggests that monitoring of

tertiary folding by fluorescence of covalently attached pyrene will be generally applicable to structured RNA mols.

IT 880146-33-8

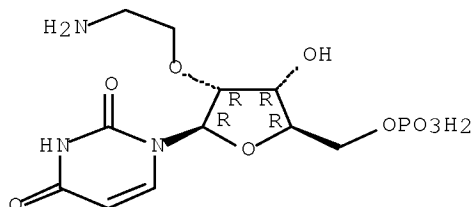
RL: BUU (Biological use, unclassified); RCT (Reactant); BIOL (Biological study); RACT (Reactant or reagent); USES (Uses)

(fluorescence of covalently attached pyrene as general RNA folding probe in relation to tether lengths and compns.)

RN 880146-33-8 CAPLUS

CN 5'-Uridylic acid, 2'-O-(2-aminoethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 46 THERE ARE 46 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 3 OF 27 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2005:480331 CAPLUS [Full-text](#)

DOCUMENT NUMBER: 143:207771

TITLE: Four base recognition by triplex-forming oligonucleotides at physiological pH

AUTHOR(S): Rusling, David A.; Powers, Vicki E. C.; Ranasinghe, Rohan T.; Wang, Yang; Osborne, Sadie D.; Brown, Tom; Fox, Keith R.

CORPORATE SOURCE: School of Biological Sciences, University of Southampton, Southampton, SO16 7PX, UK

SOURCE: Nucleic Acids Research (2005), 33(9), 3025-3032  
CODEN: NARHAD; ISSN: 0305-1048

PUBLISHER: Oxford University Press

DOCUMENT TYPE: Journal

LANGUAGE: English

AB We have achieved recognition of all 4 bp by triple helix formation at physiolo. pH, using triplex-forming oligonucleotides that contain four different synthetic nucleotides. BAU [2'-aminoethoxy-5-(3-aminoprop-1-ynyl)uridine] recognizes AT base pairs with high affinity, MeP (3-methyl-2 aminopyridine) binds to GC at higher pHs than cytosine, while APP (6-(3-aminopropyl)-7-methyl-3H-pyrrolo[2,3-d]pyrimidin-2(7H)-one) and S [N-(4-(3-acetamidophenyl)thiazol-2-yl-acetamide)] bind to CG and TA base pairs, resp. Fluorescence melting and DNase I footprinting demonstrate successful triplex formation at a 19mer oligopurine sequence that contains two CG and two TA interruptions. The complexes are pH dependent, but are still stable at pH 7.0. BAU, MeP and APP retain considerable selectivity, and single base pair changes opposite these residues cause a large reduction in affinity. In contrast, S is less selective and tolerates CG pairs as well as TA.

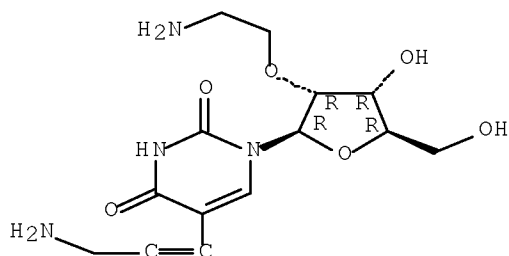
IT 861711-48-0

RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)

(base pair selectivity of synthetic nucleotide; four base recognition by triplex-forming oligonucleotides at physiolo. pH)

RN 861711-48-0 CAPLUS  
 CN Uridine, 2'-O-(2-aminoethyl)-5-(3-amino-1-propynyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 50 THERE ARE 50 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 4 OF 27 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2004:400638 CAPLUS Full-text

DOCUMENT NUMBER: 141:118826

TITLE: 2'-O-[2-(Guanidinium)ethyl]-Modified Oligonucleotides:

Stabilizing Effect on Duplex and Triplex Structures

AUTHOR(S): Prakash, Thazha P.; Pueschl, Ask P.; Lesnik, Elena; Mohan, Venkatraman; Tereshko, Valentina; Egli, Martin; Manoharan, Muthiah

CORPORATE SOURCE: Department of Medicinal Chemistry, Isis Pharmaceuticals, Inc., Carlsbad, CA, 92008, USA

SOURCE: Organic Letters (2004), 6(12), 1971-1974

CODEN: ORLEF7; ISSN: 1523-7060

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

OTHER SOURCE(S): CASREACT 141:118826

AB Oligonucleotides with a novel 2'-O-[2-(guanidinium)ethyl] (2'-O-GE) modification have been synthesized using a novel protecting group strategy for the guanidinium group. This modification enhances the binding affinity of oligonucleotides to RNA as well as duplex DNA ( $\Delta T_m$  3.2° per modification). The 2'-O-GE modified oligonucleotides exhibited exceptional resistance to nuclease degradation. The crystal structure of a palindromic duplex formed by a DNA oligonucleotide with a single 2'-O-GE modification was solved at 1.16 Å resolution.

IT 242147-79-1

RL: RCT (Reactant); RACT (Reactant or reagent)

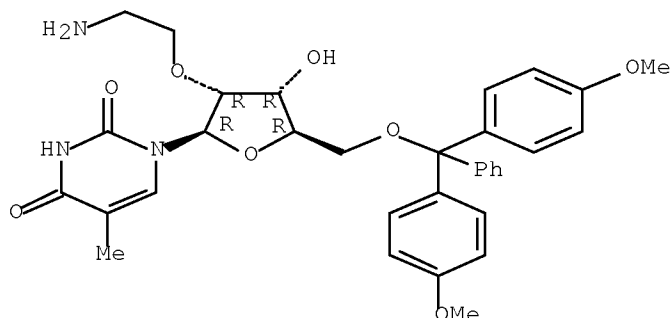
(preparation of (guanidinium)ethyl-modified oligonucleotides and interactions with dsDNA and RNA)

RN 242147-79-1 CAPLUS

CN Uridine, 2'-O-(2-aminoethyl)-5'-O-[bis(4-methoxyphenyl)phenylmethyl]-5-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.





REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 5 OF 27 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2003:217995 CAPLUS Full-text

DOCUMENT NUMBER: 138:205306

TITLE: Preparation of guanidinium functionalized oligodeoxyribonucleotide triplexes useful for diagnostic, therapeutic, and investigative purposes

INVENTOR(S): Manoharan, Muthiah; Cook, Phillip Dan; Prakash, Thazha P.; Mohan, Venkatraman

PATENT ASSIGNEE(S): Isis Pharmaceuticals, Inc., USA

SOURCE: U.S., 51 pp., Cont.-in-part of U.S. Ser. No. 349,040. CODEN: USXXAM

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

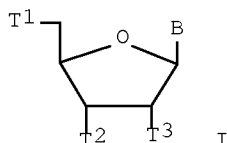
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 6534639	B1	20030318	US 2000-612531	20000707
US 6593466	B1	20030715	US 1999-349040	19990707
US 20030092046	A1	20030515	US 2002-247893	20020920
US 6914148	B2	20050705		

PRIORITY APPLN. INFO.: US 1999-349040 A2 19990707  
US 2000-612531 A3 20000707

OTHER SOURCE(S): CASREACT 138:205306; MARPAT 138:205306

GI



AB The present invention provides oligodeoxyribonucleotide triplexes I wherein B is nucleobase; T1 is OH, protected hydroxyl group; T2 is active phosphorus group, linking moiety attached to a solid support; T3 is H, OH, protected

hydroxyl group, sugar substituent, which are specifically hybridizable with a selected sequence of RNA or DNA wherein at least one of the nucleoside moieties of the oligomer is modified to include a guanidinium group. These oligomers are useful for diagnostic, therapeutic, and investigative purposes. Thus, 5'-O-DMT-2'-O-[2-(N-,N'- cyanoethoxycarbonyloxy-guanidinium)ethyl]-3-O-succinyl-5-methyluridine was prepared and incorporated into DNA triplexes.

IT 242147-79-1F

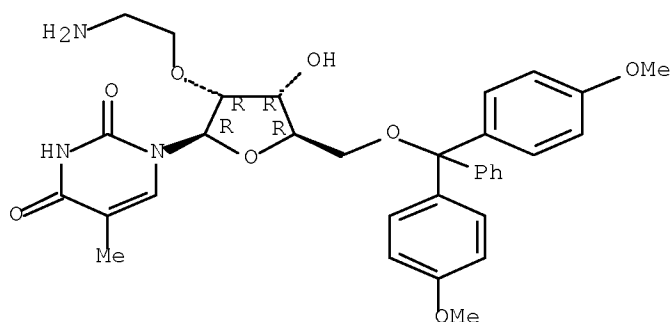
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of guanidinium functionalized oligodeoxyribonucleotide triplexes useful for diagnostic, therapeutic, and investigative purposes)

RN 242147-79-1 CAPLUS

CN Uridine, 2'-O-(2-aminoethyl)-5'-O-[bis(4-methoxyphenyl)phenylmethyl]-5-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 158 THERE ARE 158 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 6 OF 27 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2001:31524 CAPLUS Full-text

DOCUMENT NUMBER: 134:116187

TITLE: Preparation of guanidinium functionalized oligodeoxyribonucleotide triplexes

INVENTOR(S): Manoharan, Muthiah; Cook, Phillip Dan; Prakash, Thazha P.; Mohan, Venkatraman

PATENT ASSIGNEE(S): Isis Pharmaceuticals, Inc., USA

SOURCE: PCT Int. Appl., 108 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

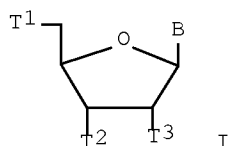
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001002423	A2	20010111	WO 2000-US18609	20000707
WO 2001002423	A3	20010503		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,				

LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,  
SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,  
YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM  
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,  
DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,  
CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG  
US 6593466 B1 20030715 US 1999-349040 19990707  
AU 2000060770 A 20010122 AU 2000-60770 20000707  
EP 1206574 A2 20020522 EP 2000-947107 20000707  
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,  
IE, SI, LT, LV, FI, RO, MK, CY, AL  
JP 2004500330 T 20040108 JP 2001-507859 20000707  
PRIORITY APPLN. INFO.: US 1999-349040 A2 19990707  
WO 2000-US18609 W 20000707  
OTHER SOURCE(S): MARPAT 134:116187  
GI



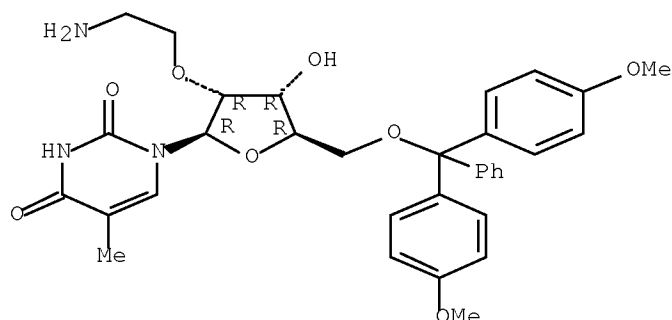
AB The present invention provides oligodeoxyribonucleotide triplexes I wherein B is nucleobase; T1 is OH, protected hydroxyl group; T2 is active phosphorus group, linking moiety attached to a solid support; T3 is H, OH, protected hydroxyl group, sugar substituent, which are specifically hybridizable with a selected sequence of RNA or DNA wherein at least one of the nucleoside moieties of the oligomer is modified to include a guanidinium group. These oligomers are useful for diagnostic, therapeutic and investigative purposes. Thus, 5'-O-DMT-2'-O-[2-(N-,N'- cyanoethoxycarbonyloxy-guanidinium)ethyl]-3-O-succinyl-5-methyluridine was prepared and incorporated into DNA triplexes. Methyluridine.

IT 242147-79-1F  
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)  
(preparation of guanidinium functionalized oligodeoxyribonucleotide triplexes)

RN 242147-79-1 CAPLUS

CN Uridine, 2'-O-(2-aminoethyl)-5'-O-[bis(4-methoxyphenyl)phenylmethyl]-5-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L13 ANSWER 7 OF 27 CAPLUS COPYRIGHT 2008 ACS on STN  
 ACCESSION NUMBER: 2000:175819 CAPLUS [Full-text](#)  
 DOCUMENT NUMBER: 132:208092  
 TITLE: Preparation of conjugated electrophilic haloacetyl  
 linked oligonucleotides  
 INVENTOR(S): Manoharan, Muthiah  
 PATENT ASSIGNEE(S): Isis Pharmaceuticals, Inc., USA  
 SOURCE: PCT Int. Appl., 65 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000014098	A1	20000316	WO 1999-US19828	19990827
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
US 6335437	B1	20020101	US 1998-149156	19980907
AU 9955899	A1	20000327	AU 1999-55899	19990827
EP 1112279	A1	20010704	EP 1999-942546	19990827
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PRIORITY APPLN. INFO.:			US 1998-149156	A1 19980907
			WO 1999-US19828	W 19990827

OTHER SOURCE(S): MARPAT 132:208092

AB The present invention provides novel methods for preparing oligonucleotide conjugates  $R_2CH_2CONH(CH_2)_n(Q)xOP(R_3)OR_1$  using a novel electrophilic haloacetyl linker, wherein  $R_1$  is a phosphorus protecting group;  $R_2$  is chlorine or a pendant group;  $R_3$  is substituted amine, heterocycloalkyl, heterocycloalkenyl;  $x$  is 0 or 1;  $n$  is 1-10;  $Q$  is a nucleoside residue. Thus, 3',5'-O-(tetraisopropylidisiloxane-1,3-diyl)-N<sup>2</sup>-(3-aminopropyl)-9-(2'-deoxy-β-D-erythro-pentafuranosyl)guanosine was prepared and incorporated into oligodeoxyribonucleotides.

IT 242147-79-1P

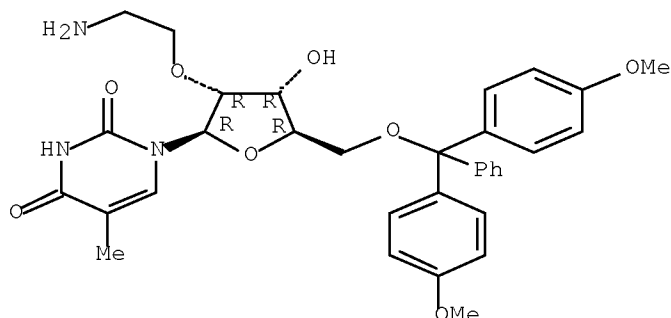
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT  
(Reactant or reagent)

(preparation of conjugated electrophilic haloacetyl linked  
oligonucleotides)

RN 242147-79-1 CAPLUS

CN Uridine, 2'-O-(2-aminoethyl)-5'-O-[bis(4-methoxyphenyl)phenylmethyl]-5-  
methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 8 OF 27 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2000:175780 CAPLUS [Full-text](#)

DOCUMENT NUMBER: 132:222803

TITLE: Preparation of Oligodeoxyribonucleotides using  
nucleobase protecting groups

INVENTOR(S): Manoharan, Muthiah

PATENT ASSIGNEE(S): Isis Pharmaceuticals, Inc., USA

SOURCE: PCT Int. Appl., 34 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

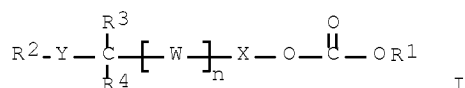
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000014048	A1	20000316	WO 1999-US19811	19990827
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
US 6166239	A	20001226	US 1998-148763	19980904
AU 9956976	A	20000327	AU 1999-56976	19990827
US 20010053855	A1	20011220	US 2000-745642	20001222
US 6576783	B2	20030610		
PRIORITY APPLN. INFO.:			US 1998-148763	A1 19980904
			WO 1999-US19811	W 19990827

OTHER SOURCE(S): MARPAT 132:222803  
GI



AB Compds. of the invention having general formula I, wherein X is aryl or a covalent bond; Y is aryl or a covalent bond; R1 is selected from succinimid-N-yl, phthalimid-N-yl, pyridin-N-yl, 4-nitophenyl, N-imidazol-1-yl, benzotriazol-2-yl, pyridin-2-yl, pentafluorophenyl, tetrafluorophenyl, triazol-N-yl, tetrazol-N-yl and norbornan-N-yl; R2 is cyano, nitro or CF3; R3 and R4 are each independently selected from the group consisting of H, alkyl, alkenyl, alkynyl, aryl, aralkyl, cycloalkyl and cycloalkylalkyl; W is C(R5)(R6) or C(R7)=C(R7) where each R5, R6, and R7 is independently selected from the group consisting of H, alkyl, alkenyl, alkynyl, aryl, aralkyl, cycloalkyl and cycloalkylalkyl or both R7 substituents together form an unsatd. aromatic ring; and n is an integer from 0 to 7 are useful reagents for protecting amine, guanidine, amidine or hydroxyl groups during organic synthesis. In particular, compds. are useful during oligonucleotide synthesis to protect nucleobase amines as well as tethered amines used for attaching functional moieties to oligonucleotides. .

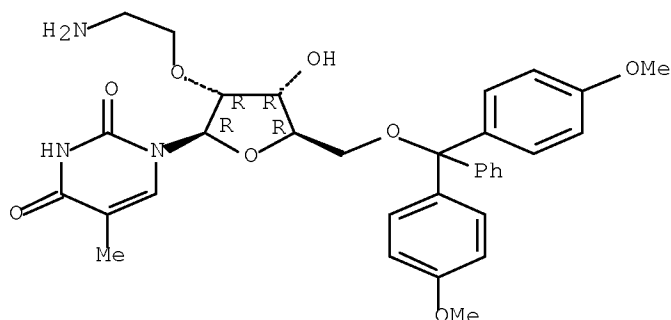
IT 242147-79-1P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)  
(preparation of oligodeoxyribonucleotides using nucleobase protecting groups)

RN 242147-79-1 CAPLUS

CN Uridine, 2'-O-(2-aminoethyl)-5'-O-[bis(4-methoxyphenyl)phenylmethyl]-5-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



IT 242147-87-1P

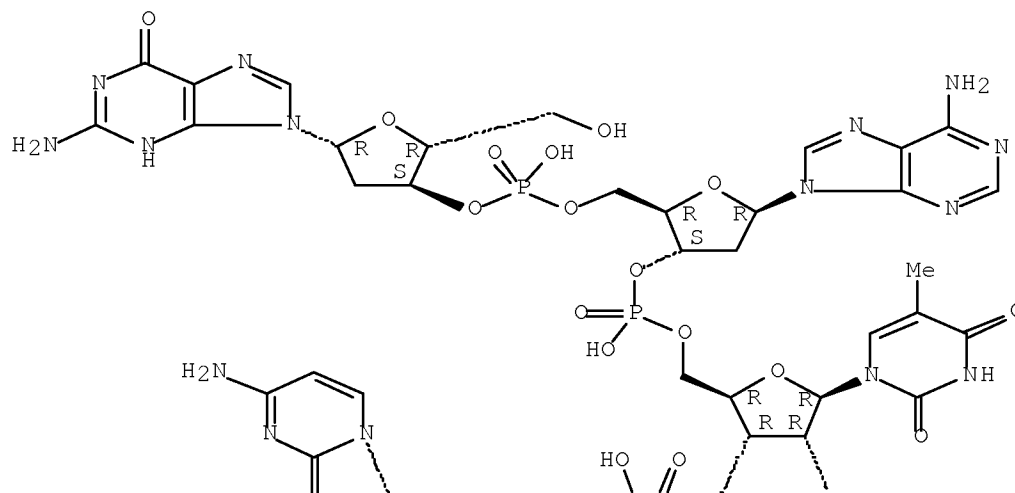
RL: SPN (Synthetic preparation); PREP (Preparation)  
(preparation of oligodeoxyribonucleotides using nucleobase protecting groups)

RN 242147-87-1 CAPLUS

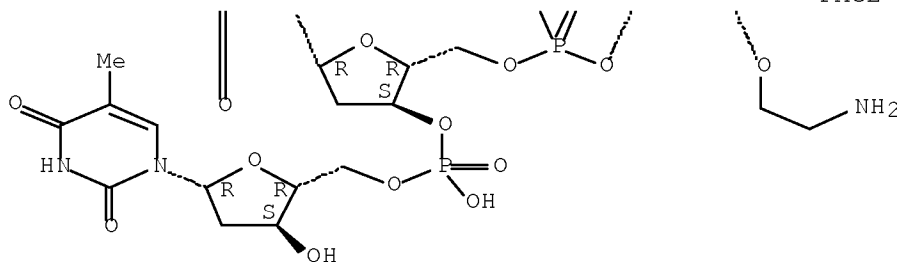
CN Thymidine, 2'-deoxyguanylyl-(3'→5')-2'-deoxyadenylyl-(3'→5')-  
2'-O-(2-aminoethyl)-5-methyluridylyl-(3'→5')-2'-deoxycytidylyl-  
(3'→5')- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A



PAGE 2-A



REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 9 OF 27 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 1999:469252 CAPLUS Full-text

DOCUMENT NUMBER: 131:228937

TITLE: N-(2-Cyanoethoxycarbonyloxy)succinimide: A New Reagent  
for Protection of Amino Groups in Oligonucleotides  
AUTHOR(S): Manoharan, Muthiah; Prakash, Thazha P.; Barber-Peoc'h,  
Isabelle; Bhat, Balkrishen; Vasquez, Guillermo; Ross,  
Bruce S.; Cook, P. Dan

CORPORATE SOURCE: Department of Medicinal Chemistry, Isis  
Pharmaceuticals, Carlsbad, CA, 92008, USA

SOURCE: Journal of Organic Chemistry (1999), 64(17), 6468-6472  
 CODEN: JOCEAH; ISSN: 0022-3263  
 PUBLISHER: American Chemical Society  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English

AB A convenient crystalline reagent to protect aminolinkers (CEOC-O-succinimide) has been developed and used to protect nucleoside-based 2'-O-alkyl amino-linkers and non-nucleosidic amino-linkers. After oligodeoxyribonucleotide synthesis incorporating these amino-linkers, standard NH<sub>4</sub>OH treatment removes the CEOC group by  $\beta$ -elimination. The resultant oligodeoxyribonucleotides modified with 2'-O-alkylamines stabilize antisense oligomers toward RNA binding. Amino-linkers generated by this new method are also useful for conjugation chemical. In addition to the potential applications in the nucleic acid field, the CEOC reagent is of general use for amino groups in all classes of compds.

IT 242147-79-1P

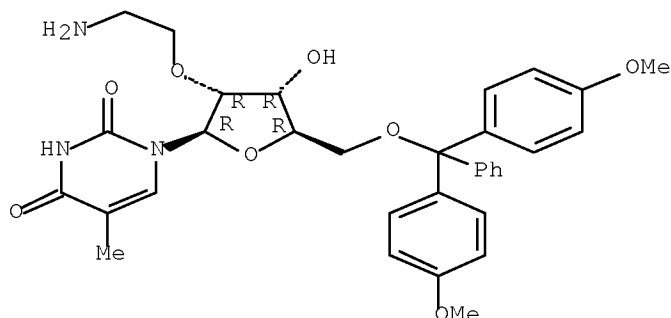
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(N-(2-cyanoethoxycarbonyloxy)succinimide as new reagent for protection of amino groups in preparation of oligodeoxyribonucleotides)

RN 242147-79-1 CAPLUS

CN Uridine, 2'-O-(2-aminoethyl)-5'-O-[bis(4-methoxyphenyl)phenylmethyl]-5-methyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



IT 242147-87-1P

RL: SPN (Synthetic preparation); PREP (Preparation)

(N-(2-cyanoethoxycarbonyloxy)succinimide as new reagent for protection of amino groups in preparation of oligodeoxyribonucleotides)

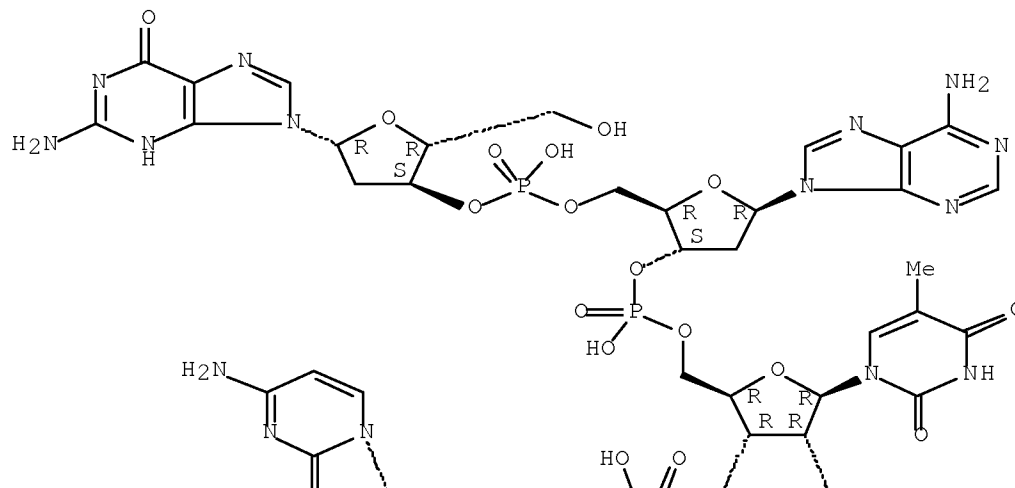
RN 242147-87-1 CAPLUS

CN Thymidine, 2'-deoxyguanylyl-(3'→5')-2'-deoxyadenylyl-(3'→5')-2'-O-(2-aminoethyl)-5-methyluridylyl-(3'→5')-2'-deoxycytidylyl-(3'→5')- (9CI) (CA INDEX NAME)

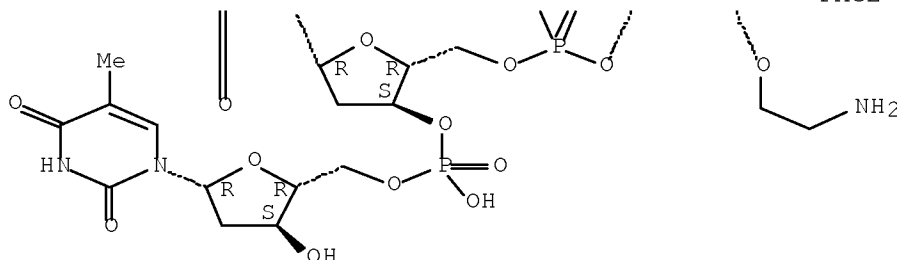
Absolute stereochemistry.



PAGE 1-A



PAGE 2-A



REFERENCE COUNT: 27 THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 10 OF 27 WPIX COPYRIGHT 2008 THOMSON REUTERS on STN  
 ACCESSION NUMBER: 2004-031184 [03] WPIX  
 CROSS REFERENCE: 2001-138119; 2003-644179; 2004-118052  
 DOC. NO. CPI: C2004-010287 [03]  
 TITLE: New oligomers containing guanidinium groups, useful for modulating gene expression by hybridizing oligomer with single- or double-stranded nucleic acids  
 DERWENT CLASS: A96; B04; D16  
 INVENTOR: COOK P D; MANOHARAN M; MOHAN V; PRAKASH T P  
 PATENT ASSIGNEE: (COOK-I) COOK P D; (ISIS-N) ISIS PHARM INC; (MANO-I) MANOHARAN M; (MOHA-I) MOHAN V; (PRAK-I) PRAKASH T P  
 COUNTRY COUNT: 1

PATENT INFO ABBR.:

PATENT NO	KIND DATE	WEEK	LA	PG	MAIN IPC
-----					

US 20030092046 A1 20030515 (200403)\* EN 54[22]  
 US 6914148 B2 20050705 (200544) EN

## APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
US 20030092046	A1 CIP of	US 1999-349040	19990707
US 20030092046	A1 Div Ex	US 2000-612531	20000707
US 20030092046	A1	US 2002-247893	20020920
US 6914148	B2 CIP of	US 1999-349040	19990707
US 6914148	B2 Div Ex	US 2000-612531	20000707
US 6914148	B2	US 2002-247893	20020920

## FILING DETAILS:

PATENT NO	KIND	PATENT NO
US 6914148	B2 Div ex	US 6534639 B
US 6914148	B2 CIP of	US 6593466 B

PRIORITY APPLN. INFO: US 2002-247893 20020920  
 US 1999-349040 19990707  
 US 2000-612531 20000707

AN 2004-031184 [03] WPIX  
 CR 2001-138119; 2003-644179; 2004-118052  
 AB US 20030092046 A1 UPAB: 20060120

NOVELTY - An oligomer comprising several nucleotide units (I), is new.

DETAILED DESCRIPTION - An oligomer comprising several nucleotide units of formula (I), is new.

B = heterocyclic base (optionally substituted by R1;

T1, T2 = OH, protected OH, nucleotide, nucleoside, or oligonucleotide;

and

T3 = H, OH, protected OH or sugar substituent group; or

T3 = at least one R1, occurring at the 3'-end or 5'-end;

R1 = a group of formula (i) or (ii);

Z = single bond, O, N or S;

R2-R4, R'3, R'4 = H, C(O)R5, or 1-10C alkyl, 2-10C alkenyl or 2-10C alkynyl (all optionally substituted by OH, NH2, alkoxy, carboxy, benzyl, phenyl, NO2, SH, thioalkoxy, halo, alkyl, aryl, alkenyl or alkynyl), alkylsulfonyl, arylsulfonyl, chemical functional group or conjugate group; or

R3+R4 = R7;

R'3+R'4 = R'7;

R5 = 1-10C alkyl, CF3, cyanoethoxy, OMe, OEt, Ot-Bu, allyloxy, 9-fluorenylmethoxy, 2-(trimethylsilyl)-ethoxy, 2,2,2-trichloroethoxy, benzyloxy, butyryl, iso-butyryl, phenyl or aryl;

R7, R'7 = H or forms a phthalimide moiety with the N atom to which it is attached;

m = 0-1; and

n = 1-6.

INDEPENDENT CLAIMS are also included for:

(1) monomer units of formula (I');

(2) preparation of (I); and

(3) compounds of formula (II)-(VI).

T'1 = T1;

T'2 = an activated phosphorous group or a linking moiety attached to a solid support;

T'3 = T3; or

T'1, T'2 and/or T'3 = R1;

CPG = controlled pore glass protecting group;

DMT = dimethoxytrityl protecting group;

X1 = cyanoethyloxy, benzyloxy, t-butoxy, methoxy, ethoxy, allyloxy, 9-fluorenylmethoxy, 2-(trimethylsilyl)-ethoxy, 2,2,2-trichloroethoxy, trifluoromethyl, butyryl, iso-butyryl, phenyl or aryl;

X2 = 3-10C alkyl, 6-24C aryl, 6-24C heteroaryl, 4-20C alicyclic, 4-20C alicyclic having at least one heteroatom, nucleoside, nucleotide or oligonucleotide;

Y1 = OH protecting group;

Y2 = activated phosphorous group or a linking moiety attached to a solid support; and

R6, R'6 = CF3, cyanoethyloxy, OMe, OEt, Ot-Bu, allyloxy, 9-fluorenylmethoxy, 2-(trimethylsilyl)-ethoxy, 2,2,2-trichloroethoxy, benzyloxy, butyryl, iso-butyryl, phenyl or aryl.

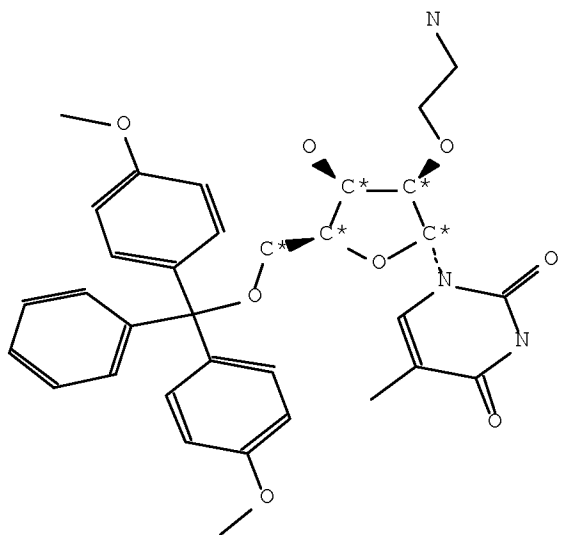
USE - The oligomers are useful for modulating gene expression which involves hybridizing with single-stranded or double-stranded nucleic acids (claimed).

DESCRIPTION OF DRAWINGS - The figure shows binding affinity of 2'-O-(2-(guanidinium)ethyl) modified oligonucleotides as a function of position in placement compared to 2'-aminopropyl and 2'-O- dimethylaminoethoxyethyl (DMAEOE).

AN.S DCR-789658

CN.S 1-{3-(2-Amino-ethoxy)-5-[bis-(4-methoxy-phenyl)-phenyl-methoxymethyl]-4-hydroxy-tetrahydro-furan-2-yl}-5-methyl-1H-pyrimidine-2,4-dione

SDCN RABXZ8



L13 ANSWER 11 OF 27 MARPAT COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 148:26120 MARPAT [Full-text](#)

TITLE: Small interfering double-stranded RNAs with chemically modified oligonucleotide chains that facilitate RISC loading for RNA interference of gene expression

INVENTOR(S): Swayze, Eric E.; Bhat, Balkrishen; Griffey, Richard  
 H.; Prakash, Thazha P.; Allerson, Charles  
 PATENT ASSIGNEE(S): Isis Pharmaceuticals, Inc., USA  
 SOURCE: U.S. Pat. Appl. Publ., 66pp., Cont.-in-part of U.S.  
 Ser. No. 946,147.  
 CODEN: USXXCO  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 47  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 20070275921	A1	20071129	US 2007-569929	20070802
US 5898031	A	19990427	US 1996-659440	19960606
US 6107094	A	20000822	US 1997-870608	19970606
EP 1600506	A2	20051130	EP 2004-78165	19970606
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
EP 1621545	A2	20060201	EP 2004-78164	19970606
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
US 20020165189	A1	20021107	US 2002-78949	20020220
US 20040147023	A1	20040729	US 2003-701265	20031104
US 20050053976	A1	20050310	US 2004-859825	20040603
WO 2005120230	A2	20051222	WO 2004-US17485	20040603
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WO 2005121368	A1	20051222	WO 2004-US17522	20040603
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US 20050130923	A1	20050616	US 2004-946147	20040920
WO 2005121370	A2	20051222	WO 2005-US19217	20050602
WO 2005121370	A3	20060526		
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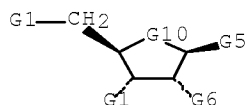
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US 20080039618 A1 20080214  
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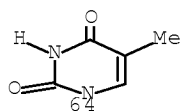
US 2007-871436 20071012  
 US 1996-659440 19960606  
 US 1997-870608 19970606  
 US 2000-479783 20000107  
 US 2002-78949 20020220  
 US 2002-423760P 20021105  
 US 2003-503271P 20030915  
 US 2003-503997P 20030918  
 US 2003-701265 20031104  
 US 2004-859825 20040603  
 WO 2004-US17485 20040603  
 WO 2004-US17522 20040603  
 US 2004-584045P 20040629  
 US 2004-607927P 20040907  
 US 2004-946147 20040920  
 WO 2005-US19217 20050602  
 EP 1997-929875 19970606  
 US 2003-701285 20031104

AB The present invention provides double-stranded compns. wherein one of the strands is useful in, for example, influencing the preferential loading the opposite strand into the RISC (or cleavage) complex. In particular, the present invention provides oligomeric compds. that comprise chemical modifications in at least one of the strands to drive loading of the opposite strand into the RISC (or cleavage) complex. Such modifications can be used to increase potency of duplex constructs that have been modified to enhance stability. Examples of chemical modifications that drive loading of the second strand include, but are not limited to, MOE (2'-O(CH<sub>2</sub>)<sub>2</sub>OCH<sub>3</sub>), 2'-O-Me, -Et, -Pr, and -N-methylacetamide. Such modifications can be distributed throughout the strand, or placed at the 5' and/or 3' ends to make a gapmer motif on the sense strand. The activity of the 4'-thio gapmer RNA antisense strand can be improved by incorporating alternating MOE or MOE gapmer motif into the sense strand.

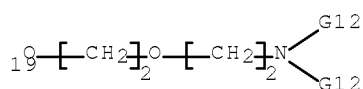
MSTR 1



G5 = 64



G6 = 19



G10 = 0

Patent location: claim 1

L13 ANSWER 12 OF 27 MARPAT COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 144:81963 MARPAT Full-text

TITLE: Double-stranded RNA compounds which facilitate RISC loading

INVENTOR(S): Swayze, Eric E.; Bhat, Balkrishen; Griffey, Richard H.; Prakash, Thazha P.; Allerson, Charles

PATENT ASSIGNEE(S): Isis Pharmaceuticals, Inc., USA

SOURCE: PCT Int. Appl., 127 pp., which which which  
CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 47

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005121370	A2	20051222	WO 2005-US19217	20050602
WO 2005121370	A3	20060526		
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
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US 20050053976	A1	20050310	US 2004-859825	20040603
WO 2005120230	A2	20051222	WO 2004-US17485	20040603
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
RW:	BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
WO 2005121368	A1	20051222	WO 2004-US17522	20040603
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LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,  
 NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,  
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 SN, TD, TG

US 20050130923 A1 20050616 US 2004-946147 20040920

EP 1765415 A2 20070328 EP 2005-756325 20050602

R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,  
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US 20070275921 A1 20071129 US 2007-569929 20070802

PRIORITY APPLN. INFO.:

US 2004-859825 20040603

WO 2004-US17485 20040603

WO 2004-US17522 20040603

US 2004-584045P 20040629

US 2004-607927P 20040907

US 2004-946147 20040920

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US 2002-78949 20020220

US 2002-423760P 20021105

US 2003-503271P 20030915

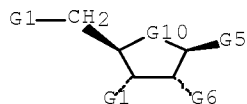
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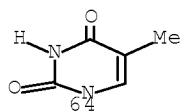
WO 2005-US19217 20050602

AB The present invention provides double stranded compns. wherein one of the strands is useful in, for example, influencing the preferential loading of the opposite strand into the RISC (or cleavage) complex. In particular, the present invention provides oligomeric compds. that comprise chemical modifications in at least one of the strands to drive loading of the opposite strand into the RISC (or cleavage) complex. Such modifications can be used to increase potency of duplex constructs that have been modified to enhance stability. Examples of chemical modifications that drive loading of the second strand include, but are not limited to, MOE (2'-O(CH<sub>2</sub>)<sub>2</sub>OCH<sub>3</sub>), 2'-O-Me, -Et, -Pr, and -N-methylacetamide. Such modifications can be distributed throughout the strand, or placed at the 5' and/or 3' ends to make a gapmer motif on the sense strand. The activity of the 4'-thio gapmer RNA antisense strand can be improved by incorporating alternating MOE or MOE gapmer motif into the sense strand.

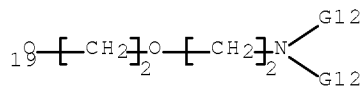
MSTR 1



G5 = 64



G6 = 19



G10 = O

Patent location: claim 1

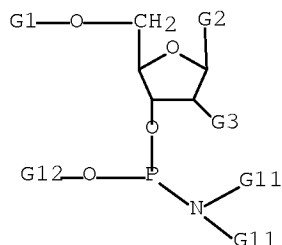
L13 ANSWER 13 OF 27 MARPAT COPYRIGHT 2008 ACS on STN  
 ACCESSION NUMBER: 141:49377 MARPAT Full-text  
 TITLE: Process for nano-filtration membrane purifying  
 oligonucleotide synthons  
 INVENTOR(S): McCormac, Paul; Hargreaves, Stephen  
 PATENT ASSIGNEE(S): Avecia Limited, UK  
 SOURCE: PCT Int. Appl., 9 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004055037	A2	20040701	WO 2003-GB5474	20031216
WO 2004055037	A3	20040916		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
CA 2510483	A1	20040701	CA 2003-2510483	20031216
AU 2003288553	A1	20040709	AU 2003-288553	20031216
EP 1590361	A2	20051102	EP 2003-780394	20031216
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK				
CN 1726223	A	20060125	CN 2003-80106420	20031216
JP 2006512336	T	20060413	JP 2004-559907	20031216
IN 2005DN02791	A	20061229	IN 2005-DN2791	20050623
US 20060135760	A1	20060622	US 2006-539202	20060126
PRIORITY APPLN. INFO.:				
			GB 2002-29423	20021218
			WO 2003-GB5474	20031216

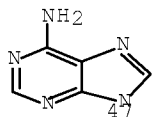


AB A process for the purification of an oligonucleotide synthon is provided. The process comprises subjecting an organic solution comprising an oligonucleotide synthon and lower mol. weight impurities to nanofiltration whereby the ratio of an oligodeoxyribonucleotide synthon to lower mol. weight impurities in the solution is increased after the nano-filtration. Preferably, the oligonucleotide synthon is a nucleoside phosphoramidite or nucleoside H-phosphonate. The nano-filtration membrane STARMEM is preferably a polyimide membrane having a mol. weight cut off of 400.

MSTR 1



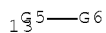
G2 = 47



G3 = 11



G4 = 13



G5 = (1-6) CH2

G6 = NH2

Patent location: claim 3

L13 ANSWER 14 OF 27 MARPAT COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 140:105281 MARPAT [Full-text](#)

TITLE: 2'-O-Modified ribosyl nucleosides and methods of enhancing renal uptake of oligonucleotides

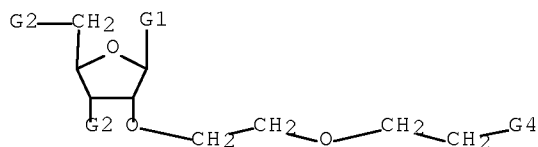
INVENTOR(S): Manoharan, Muthiah; Cook, Phillip Dan

PATENT ASSIGNEE(S): USA  
 SOURCE: U.S. Pat. Appl. Publ., 21 pp., Cont.-in-part of U.S.  
 6,600,032.  
 CODEN: USXXCO  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 3  
 PATENT INFORMATION:

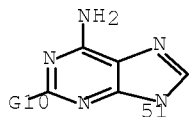
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 20040009938	A1	20040115	US 2003-359328	20030206
US 6043352	A	20000328	US 1998-130566	19980807
US 6600032	B1	20030729	US 1999-370625	19990806
PRIORITY APPLN. INFO.:			US 1998-130566	19980807
			US 1999-370625	19990806

AB 2'-O-Modified ribosyl nucleosides and modified methods containing such nucleosidic monomers are disclosed. Methods are disclosed that have increased binding affinity as shown by mol. modeling expts. Methods are also disclosed for enhancing the renal uptake of oligomeric compds. as shown using a two-step HRP imunohistochem. assay. The 2'-O-modified nucleosides of the invention include ring structures that position the sugar moiety of the nucleosides preferentially in 3' endo geometries.

MSTR 1



G1 = 51



G4 = 20



Patent location: claim 1

L13 ANSWER 15 OF 27 MARPAT COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 140:106457 MARPAT Full-text

TITLE: Synthesis of mutagen-coupled modified oligonucleotides and their use in targeted DNA modification of S phase-synchronized animal cells

INVENTOR(S): Seidman, Michael M.; Puri, Nitin; Majumdar, Alokes

PATENT ASSIGNEE(S): Dept. of Health &amp; Human Services, USA

SOURCE: U.S. Pat. Appl. Publ., 59 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: English

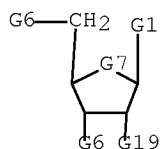
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

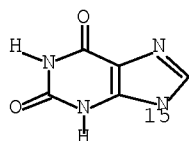
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 20040009602	A1	20040115	US 2003-438076	20030513
PRIORITY APPLN. INFO.:			US 2002-378025P	20020513

AB This invention relates to the synthesis of mutagen-coupled modified oligonucleotides and their use in targeted DNA modification of S phase-synchronized fertilized eggs. The precursor of modified oligonucleotides, 2'-AE-5-methyluridine controlled pore glass (CPG), was prepared by addition of 4-(Dimethylamino)pyridine (0.64 mmol) to 5'-O-(4,4'-Dimethoxytrityl)-5-methyluridine-2'-O-(2-aminoethoxy) (1) (0.643 mmol) over 1 h. This yielded 5'-O-(4,4'-Dimethoxytrityl)-5-methyluridine-2'-O-(2-aminoethyl)-3'-O-Succinate (2) (85% yield). Reaction of compound (2) with N,N-diisopropylethylamine (0.5 mmol), O-(7-azabenzotriazol-1 yl)-1,1,2,2-tetramethyluronium hexafluorophosphate (0.2 mmol), and finally with acid treated long chain alkylamine-CPG (90  $\mu$ mol/g) for 16 h, yielded 5'-O-(4,4'-dimethoxytrityl)-5-methyluridine-2'-O-methyl-3'-O-succinimido-N6-hexanamido-N3-propyl-controlled pore glass (3) (30  $\mu$ mol/g). Compound (3) was then used in the synthesis of modified oligonucleotides, including triplex forming oligonucleotides, peptide nucleic acids and polyamides. The primary benefit of mutagen-coupled modified oligonucleotides is the sequence specific targeting of the oligonucleotide to the target sequence in the cell genome, directing the mutagen to the desired site of mutagenesis. Further, introduction of the mutagen during S phase induces a higher rate of mutagenesis, compared to other cell cycle phases. Examples shown include use of these mutagens in targeted mutagenesis of the HPRT (hypoxanthine phosphoribosyl transferase) gene in animal cells. Targeted mutagenesis has the potential to be applied towards gene therapy of human diseases, by alteration of mutant genes in human cells and incorporation of such cells into a patient.

MSTR 1



G1 = 15



G7 = O  
 G17 = CH<sub>2</sub>CH<sub>2</sub>  
 G19 = 170 / 253 / 256 / 259

$\text{G}^{20}_{20}-\text{G}^{17}_{17}-\text{G}^{21}_{21}$        $\text{G}^{22}_{22}-\text{G}^{24}_{24}-\text{G}^{21}_{21}$        $\text{G}^{23}_{23}-\text{G}^{26}_{26}-\text{G}^{21}_{21}$        $\text{G}^{22}_{22}-\text{G}^{25}_{25}-\text{G}^{21}_{21}$

G20 = O  
 G21 = 250



Patent location: claim 1

L13 ANSWER 16 OF 27 MARPAT COPYRIGHT 2008 ACS on STN  
 ACCESSION NUMBER: 138:90027 MARPAT [Full-text](#)  
 TITLE: Preparation of oligodeoxyribonucleotides via  
 condensation reaction using 1,1-dioxo-1,2-dihydro-  
 1λ6-benzo[d]isothiazol-3-one salt activators  
 INVENTOR(S): Sinha, Nanda; Zedalis, William Edward; Miranda,  
 Gregory Keith  
 PATENT ASSIGNEE(S): Avecia Biotechnology Inc., USA; Avecia Limited  
 SOURCE: PCT Int. Appl., 43 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 4  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2003004512	A1	20030116	WO 2002-GB3029	20020701
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CA 2452205	A1	20030116	CA 2002-2452205	20020701
AU 2002319409	A1	20030121	AU 2002-319409	20020701
EP 1404696	A1	20040407	EP 2002-748994	20020701
EP 1404696	B1	20060208		

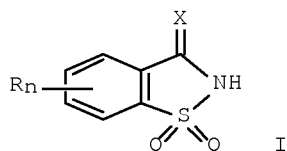
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JP 2004533488	T	20041104	JP 2003-510678	20020701
CN 1549820	A	20041124	CN 2002-817156	20020701
AT 317395	T	20060215	AT 2002-748994	20020701
ES 2258151	T3	20060816	ES 2002-748994	20020701
HU 2004000151	A2	20070828	HU 2004-151	20020701
IN 2003DN02244	A	20060120	IN 2003-DN2244	20031223
US 20060041114	A1	20060223	US 2004-482441	20040813
IN 2005DN02792	A	20061229	IN 2005-DN2792	20050623

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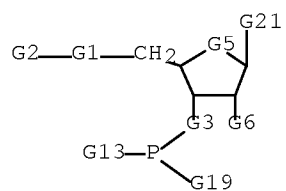
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WO 2002-GB3029	20020701
GB 2002-29443	20021218

GI

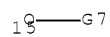


AB A process for the synthesis of oligonucleotides using phosphoramidite chemical is provided. The process employs as activator a 1,1-dioxo-1,2-dihydro-1λ6-benzo[d]isothiazol-3-one, preferably in the presence of an organic base. The 1,1-dioxo-1,2-dihydro-1λ6-benzo[d]isothiazol-3-one is represented by the following structural formula I; wherein n is 0 or an integer from 1 to 4; X is O or S; R for each occurrence is a substituent, preferably each independently, a halo, a substituted or unsubstituted aliphatic group, -NR<sub>1</sub>R<sub>2</sub>, -OR<sub>3</sub>, -OC(O)R<sub>3</sub>, -C(O)OR<sub>3</sub>, or cyano; or two adjacent R groups taken together with the carbon atoms to which they are attached form a six membered saturated or unsatd. ring; R<sub>1</sub> and R<sub>2</sub> are each, independently, H, a substituted or unsubstituted aliphatic group, a substituted or unsubstituted aryl group, or a substituted or unsubstituted aralkyl group; and R<sub>3</sub> is a substituted or unsubstituted aliphatic group, a substituted or unsubstituted aryl group, or a substituted or unsubstituted aralkyl group. Preferred organic bases are pyridine, 3-methylpyridine, or N-methylimidazole. Thus, 5'-TCTCCCAGCGTGCGCCAT-3' was prepared via condensation reaction using salt activator 1,1-dioxo-1,2-dihydro-1λ6-benzo[d]isothiazol-3-one and N-methylimidazole.

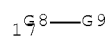
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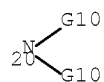
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G6 = 15



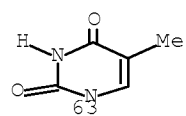
G7 = 17



G8 = (1-6) CH2  
G9 = 20

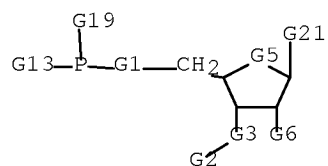


G21 = 63



Patent location: claim 32

MSTR 3




G5 = 0  
G6 = 15

~~18~~—G7

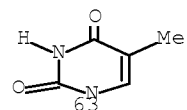
G7 = 17

~~19~~—G8—G9

G8 = (1-6) CH<sub>2</sub>  
G9 = 20

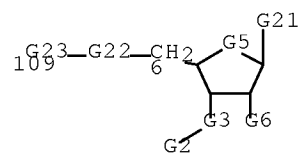
~~20~~ 

G21 = 63



Patent location: claim 32

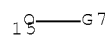
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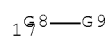
G2 = 110

~~110~~—G8—G9

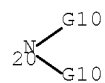
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G6 = 15



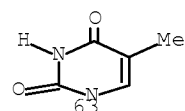
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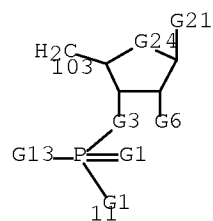
G8 = (1-6) CH2  
G9 = 20



G21 = 63



G22 = (0-3) 103-109 11-6

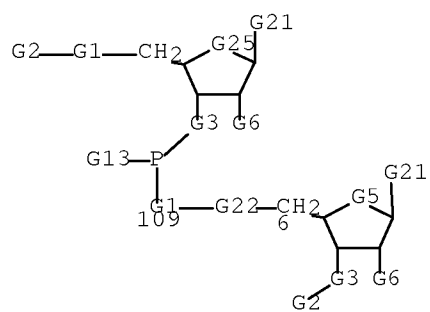


Patent location:  
Note:

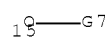
claim 35  
oligonucleotides containing additional nucleotide  
units also claimed

MSTR 5

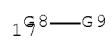




G6 = 15

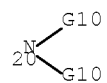


G7 = 17

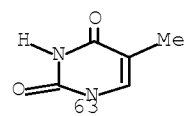


G8 = (1-6) CH2

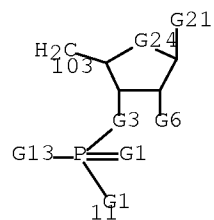
G9 = 20



G21 = 63



G22 = (0-3) 103-109 11-6



G24 = 0

Patent location:

claim 35

Note:

oligonucleotides containing additional nucleotide  
units also claimed

REFERENCE COUNT:

11

THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 17 OF 27 MARPAT COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 139:96329 MARPAT Full-textTITLE: 2'-O-Aminoethyloxyethyl-modified oligonucleotides and  
their use for inhibiting gene expression

INVENTOR(S): Manoharan, Muthiah; Cook, Phillip Dan

PATENT ASSIGNEE(S): ISIS Pharmaceuticals, Inc., USA

SOURCE: U.S., 26 pp., Cont.-in-part of U.S. 6,043,352.

CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

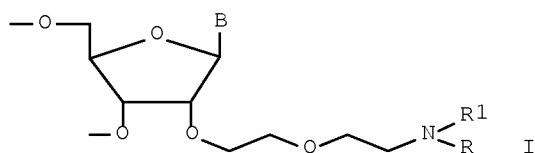
English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

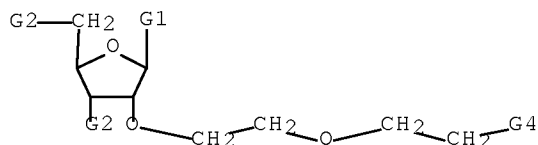
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 6600032	B1	20030729	US 1999-370625	19990806
US 6043352	A	20000328	US 1998-130566	19980807
US 6673912	B1	20040106	US 2002-121135	20020411
US 20040009938	A1	20040115	US 2003-359328	20030206
PRIORITY APPLN. INFO.:			US 1998-130566	19980807
			US 1999-370625	19990806

GI

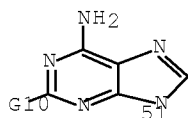


AB 2'-O-Modified ribosyl nucleosides and modified oligomeric compds. containing such nucleosidic monomers I, wherein R and R1 are independently H, nitrogen protecting group, alkyl, alkenyl, alkynyl; R and R1 together are nitrogen protecting group, joined in N- and O-containing heterocycle, are disclosed. Oligomeric compds. are disclosed that have increased binding affinity as shown by mol. modeling expts. The 2'-O-modified nucleosides of the invention include ring structures that position the sugar moiety of the nucleosides preferentially in 3' endo geometries.

MSTR 1



G1 = 51



G4 = 20



Patent location: claim 1

REFERENCE COUNT: 114 THERE ARE 114 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE REFORMAT

L13 ANSWER 18 OF 27 MARPAT COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 137:268427 MARPAT [Full-text](#)

TITLE: Antitumoral, antiviral, antibacterial, antiparasitic, anti-inflammatory, immunomodulatory and antimycotic medicinal preparation, method for production and dosage forms thereof

INVENTOR(S): Travkin, Oleg Viktorovich; Yakovleva, Elena Vladimirovna

PATENT ASSIGNEE(S): Russia

SOURCE: PCT Int. Appl., 28 pp.  
CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: Russian

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002074311	A1	20020926	WO 2002-RU117	20020318
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, RU				

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH,  
 CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR,  
 BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

RU 2197248 C2 20030127 RU 2001-107818 20010320

AU 2002308901 A1 20021003 AU 2002-308901 20020318

PRIORITY APPLN. INFO.:

RU 2001-107818 20010320

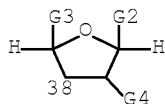
WO 2002-RU117 20020318

AB The invention relates to medicine, pharmacol. and to the chemical and pharmaceutical industry. The aim of the invention is to develop a new medicinal preparation having high activity and having a large range of the above mentioned non-specific pharmacol. action, and a production method for all presently known dosage forms. The novelty of the invention lies in a biol. active part of the preparation which contains a mixture of acridoneacetic acid or the derivs. thereof with monosubstituted ethers of monosaccharides, the biol. active part being heat-treated together with a pharmaceutically acceptable carrier at temps. ranging from 80 °C to 120 °C. The medicinal preparation containing ANANDIN exhibits higher efficiency. ANANDIN is embodied in the form of a mixture of N-acridoneacetic acid (glucoaminopropylcarbaccridone) and 3-O-(N,N-dimethylamino-n-propyl)-1.2:5.6-di-O-isopropylidene- $\alpha$ ,D- glucofuranose (dimethylaminopropylglucofuranose). The inventive medicinal preparation is embodied in the form of all known dosage forms. Six series of examples for producing the preparation and the dosage forms thereof, detailed information on the exptl. estimate of all biol. activities enumerated in the title provided with 7 tables of comparative measuring are also disclosed. The results demonstrate that a new, easily produced, stable, medicinal preparation usable for all dosage forms is created. The biol. activity of said preparation exceeds the biol. activity of analogous prepns., and the range of pharmacol. action thereof is non-specific and unusually large.

MSTR 2

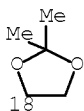
G1—G5—G6

G1 = 38



G2 = CH2Ph

G3 = 18



G5 = G8  
G6 = NH2  
G8 = (1-6) CH2

Patent location: claim 1  
Note: and hemiacetal or hemiketal forms  
Note: substitution is restricted

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 19 OF 27 MARPAT COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 135:195751 MARPAT Full-text

TITLE: Preparation of DNA having improved hybridization  
affinity and nuclease resistance via alkylation of  
nucleosides with cyclic sulfates

INVENTOR(S): Fraser, Allister S.; Manoharan, Muthiah; Cook, Phillip  
Dan; Jung, Michael E.; Kawasaki, Andrew M.

PATENT ASSIGNEE(S): Isis Pharmaceuticals, Inc., USA

SOURCE: U.S., 45 pp.  
CODEN: USXXAM

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

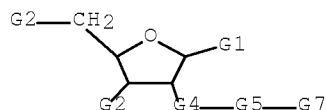
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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US 6277982	B1	20010821	US 1999-378665	19990820
PRIORITY APPLN. INFO.:			US 1999-378665	19990820
OTHER SOURCE(S):		CASREACT 135:195751		

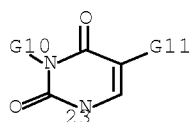
GI

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

AB Methods for the alkylation of alcs., amines and thiols by the use of cyclic sulfates are disclosed. The alkylated sulfates formed are versatile intermediates which may be further elaborated by methods of the invention. In particular, methods for the alkylation of the 2', 3' or 5'-hydroxy position of nucleosides and nucleoside analogs with cyclic sulfates to form the 2', 3' or 5'-O-alkyl sulfate modified compds. are disclosed. Displacement of the 2',3' or 5'-O-sulfate with a nucleophile provides 2', 3' or 5'-O-modified nucleosides and nucleoside analogs useful for the synthesis of oligodeoxyribonucleotide compds. having improved hybridization affinity and nuclease resistance. A process for preparing a compound AX(CR1R2)nOSO2O-Y+ (I) wherein: A is a carbohydrate, oligonucleotide, nucleotide, or nucleoside; X is a O, S, or N; R1 and R2 are independently H, alkyl, aryl, O-alkyl, O-aryl, carboxylic acid, amide, ester, halogen, trifluoromethyl, or amine; n is 2-10; and, Y is H, Li, Na, K, Cs or an amine; comprising the steps of: treating a compound of formula A-X-H with cyclic sulfate II [QM is (CR1R2)n] to give I. Thus, alkylation of nucleoside III (R3 = H) with II (QM = CH2CH2) gave III (R3 = CH2CH2OSO3Na) which was used in the preparation of DNA duplexes as nuclease resistant and antiviral agent (no data).



G1 = 23



G4 = O  
 G5 = G9  
 G7 = NH2 (opt. substd.)  
 G9 = (2-3) CH2 (opt. substd.)  
 Patent location: claim 50

REFERENCE COUNT: 107 THERE ARE 107 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE REFORMAT

L13 ANSWER 20 OF 27 MARPAT COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 134:37974 MARPAT Full-text

TITLE: RNA polynucleotide modified at the ribose 2'-OH position that are still capable of acting as templates in polymerization reactions

INVENTOR(S): Goldsborough, Andrew Simon

PATENT ASSIGNEE(S): Cyclops Genome Sciences Ltd., UK

SOURCE: PCT Int. Appl., 187 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000075306	A2	20001214	WO 2000-GB1670	20000502
WO 2000075306	A3	20010517		
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW			
RW:	GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
WO 2001094626	A1	20011213	WO 2000-GB1683	20000502
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,			

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 SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW  
 RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,  
 DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,  
 CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

EP 1196631 A1 20020417 EP 2000-929665 20000502

EP 1196631 B1 20061206

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,  
 IE, SI, LT, LV, FI, RO, CY

AT 347616 T 20061215 AT 2000-929665 20000502

US 20030039985 A1 20030227 US 2001-11495 20011026

US 6867290 B2 20050315

US 20050272679 A1 20051208 US 2005-57808 20050214

PRIORITY APPLN. INFO.:

GB 1999-10154 19990430

GB 1999-10156 19990430

GB 1999-10157 19990430

GB 1999-10158 19990430

WO 2000-GB1670 20000502

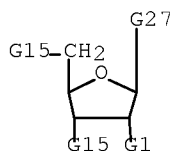
WO 2000-GB1683 20000502

WO 2000-GB1687 20000502

US 2001-11495 20011026

AB Provided is a polynucleotide comprising mRNA, rRNA or viral RNA, greater than 25% of the ribose rings of which are covalently modified at the 2'-OH position. Further provided is a method for producing a double-stranded oligo- or polynucleotide from a template, which comprises contacting the template with a plurality of mononucleotides comprising UTP, dTTP and/or dUTP, ATP and/or dATP, GTP and/or dGTP, and CTP and/or dCTP, in the presence of a nucleic acid polymerase and optionally a template primer under conditions to polymerize the mononucleotides to form a nucleic acid strand complementary to the template, wherein the template comprises an oligo- or polyribonucleotide, a proportion of the ribose rings of which are covalently modified at the 2' - OH position to bear a substituent which enables replication of the template by the nucleic acid polymerase. Thus, for example, RNA may be modified by formylation at the ribose 2'-OH position and still serve as an excellent template for reverse transcriptases. The consequence of modifying the ribose 2'-OH groups is to increase the stability and intactness of the RNA, allowing complete cDNA copies and accurate measurements of its size and abundance to be made. It is preferable to choose 2'-OH modifications that provide the maximum stability to the modified RNA, yet can be readily removed under mild conditions without leading to RNA chain cleavage. A wide variety of acylation, acetylation, and other modification reaction reagents and catalysts are described. Also provided is use of a polynucleotide comprising mRNA, rRNA or viral RNA, a proportion of the ribose rings of which are covalently modified at the 2' -OH position, in a hybridization reaction. RNA modified by acetylation has altered hybridization properties, probably reflecting a lower T<sub>m</sub> of the hybrid, and standard conditions for Northern blotting are probably too stringent and a lower temperature should be chosen.

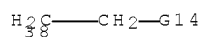
MSTR 1



G1 = 36

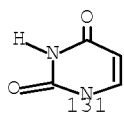


G2 = 38



G14 = NH2

G27 = 131



Patent location: claim 1

L13 ANSWER 21 OF 27 MARPAT COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 133:344607 MARPAT Full-text

TITLE: Bispecific antisense oligonucleotides inhibiting both bcl-2 and bcl-xL expression for tumor cell apoptosis induction

INVENTOR(S): Zangemeister-Wittke, Uwe; Ludke, Gerd; Huesken, Dieter

PATENT ASSIGNEE(S): Universitat Zurich, Switz.

SOURCE: PCT Int. Appl., 38 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000066724	A2	20001109	WO 2000-EP3708	20000426
WO 2000066724	A3	20010208		
W: AE, AL, AU, BA, BG, BR, CA, CN, CZ, EE, GE, HR, HU, ID, IL, IN, JP, KR, LT, LV, MK, MX, NO, NZ, PL, RO, SG, SI, SK, TR, UA, US, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
EP 1181361	A2	20020227	EP 2000-925239	20000426
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				

PRIORITY APPLN. INFO.: GB 1999-10119 19990430

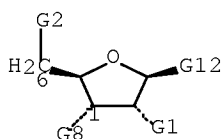
WO 2000-EP3708 20000426

AB The present invention relates to antisense oligonucleotide derivs. directed against human bcl-xL mRNA and being capable of modulating the biosynthesis of

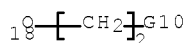


human bcl-xL protein. Furthermore, the present invention relates to antisense oligonucleotide derivs. directed against both human bcl-xL mRNA and human bcl-2 mRNA, and being capable of modulating the biosynthesis of both human bcl-xL protein and human bcl-2 protein. The present invention further relates to a pharmaceutical composition comprising such oligonucleotide derivs., uses thereof and methods of treatment and diagnosis utilizing such oligonucleotide derivs. Bcl-2 and bcl-xL are inhibitors of apoptosis frequently overexpressed in solid tumors. The bcl-2 and bcl-xL mRNAs share a region of homol. comprising nucleotides 605-624 and 687-706, resp., which differs by only 3 nucleotides. This sequence does not occur in the proapoptotic splice variant bcl-xS. To test the possibility that oligonucleotides targeting this region have the potential to down-regulate bcl-2 and bcl-xL expression simultaneously, 3 2'-O-methoxy-ethoxy-modified phosphorothioate oligonucleotides were designed. These oligonucleotides differed in the number of mismatches to bcl-2 and bcl-xL and in the number of nucleotides to which the modifications were made. The effects of these oligonucleotides on bcl-2 and bcl-xL expression, as well as their abilities to induce apoptosis, were assessed in small cell and non-small cell lung cancer cell lines expressing different basal levels of bcl-2 and bcl-xL. Although all oligonucleotides down-regulated bcl-2 and bcl-xL expression, oligonucleotide 4625, which has no mismatching nucleotides to bcl-2 but 3 to bcl-xL, 2 of which were modified by 2'-O-methoxy-ethoxy residues, showed the strongest bispecific activity on the transcript and protein level. In all cell lines this bispecific activity induced apoptotic cell death, as demonstrated by increased uptake of propidium iodide, a 10-100-fold increase in caspase-3-like protease activity, and nuclear condensation, and fragmentation. This is the 1st report of a bcl-2/bcl-xL bispecific antisense oligonucleotide that deserves attention as a therapeutic compound in lung cancer and other malignancies in which bcl-2 and/or bcl-xL are overexpressed.

MSR 1

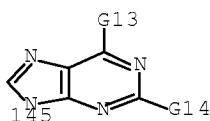


G1 = 18



G10 = NH2

G12 = 145

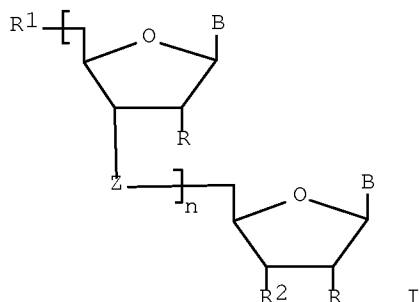


Patent location: claim 8  
 Note: substitution is restricted

L13 ANSWER 22 OF 27 MARPAT COPYRIGHT 2008 ACS on STN  
 ACCESSION NUMBER: 133:164272 MARPAT Full-text  
 TITLE: Preparation of mixed backbone antisense  
 oligodeoxyribonucleotides as inhibitors of c-raf mRNA  
 synthesis  
 INVENTOR(S): Manoharan, Muthiah; Maier, Martin A.  
 PATENT ASSIGNEE(S): Isis Pharmaceuticals, Inc., USA  
 SOURCE: PCT Int. Appl., 53 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

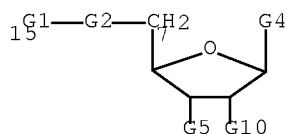
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000047593	A1	20000817	WO 2000-US3543	20000211
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
US 6207819	B1	20010327	US 1999-250075	19990212
EP 1159282	A1	20011205	EP 2000-908597	20000211
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
JP 2002536453	T	20021029	JP 2000-598512	20000211
US 20010016652	A1	20010823	US 2000-726096	20001129
US 6462184	B2	20021008		
US 20030045698	A1	20030306	US 2002-117267	20020405
PRIORITY APPLN. INFO.:			US 1999-250075	19990212
			WO 2000-US3543	20000211
			US 2000-726096	20001129

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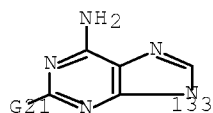


AB Synthetic processes are provided wherein mixed backbone oligomeric compds. I (B = heterocycle; Z = independently phosphodiester, phosphoramidate, boranophosphate; R = independently H, OH, protected OH; R1, R2 = independently OH, protected OH; n > 1) are prepared having at least one phosphodiester internucleoside linkage in addition to one or more phosphorothioate, phosphoramidate and boranophosphate internucleoside linkages. Novel H-phosphonate intermediates are also disclosed that are useful in the synthetic processes. The synthetic processes use a novel oxidation step to oxidize H-phosphonate internucleoside linkages to phosphodiester internucleoside linkages without degradation of adjacent phosphorothioate, phosphoramidate and boranophosphate internucleoside linkages. Thus, 5'-O-DMT-2'-MOE-N-isobutyrylguanosine-3'-H-phosphonate was prepared and incorporated into antisense oligodeoxyribonucleotide as inhibitor of c-raf mRNA synthesis. 2-(1H-benzotriazole-1-yl)-1,1,3,3- tetramethyluronium hexafluorophosphate, O-(azabenzotriazol-1-yl)-1,1,3,3- tetra-Me uronium hexafluorophosphate, 6-(trifluoromethyl)benzotriazol-1-yl- oxy-tris-pyrrolidino-phosphonium hexafluorophosphate, bromo-tris- pyrrolidino-phosphonium hexafluorophosphate, benzotriazole-1-yl-oxy-tris- pyrrolidino-phosphonium hexafluorophosphate or 2-(benzotriazol-1-yloxy)- 1,3-dimethyl-2-pyrrolidin-1-yl-1,3,2-diazaphospholidinium hexafluorophosphate.

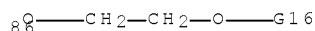
MSTR 1



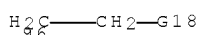
G4 = 133



G10 = 86



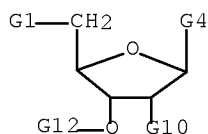
G16 = 96



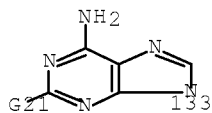
G18 = NH<sub>2</sub>

Patent location: claim 1

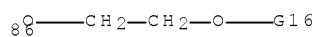
MSTR 2



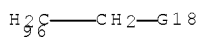
G4 = 133



G10 = 86



G16 = 96

G18 = NH<sub>2</sub>

Patent location: claim 1

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 23 OF 27 MARPAT COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 132:251377 MARPAT Full-text

TITLE: Preparation of 3',5'-cyclic phosphate nucleotides directed to bio-catalysts having cyclic phosphodiesterase activity

INVENTOR(S): Ghisalba, Oreste; Marais, Guy Joel Christian; Martin, Pierre

PATENT ASSIGNEE(S): Isis Pharmaceuticals, Inc., USA; Novartis Ag

SOURCE: PCT Int. Appl., 39 pp.

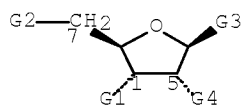
CODEN: PIXXD2

DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

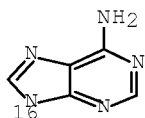
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000020429	A1	20000413	WO 1999-US23244	19991005
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
US 6147211	A	20001114	US 1998-167191	19981006
AU 9962920	A1	20000426	AU 1999-62920	19991005
US 6340754	B1	20020122	US 2000-664596	20000918
PRIORITY APPLN. INFO.:			US 1998-167191	19981006
			WO 1999-US23244	19991005

AB The present invention provides 3',5'-cyclic phosphate nucleotide compds. The present invention is further directed to processes for cleaving the cyclic phosphate moiety of the 3',5'-cyclic phosphate compds. of the present invention by treating the compds. with a biocatalyst. The present invention is also directed to bio-catalysts having cyclic phosphodiesterase activity.

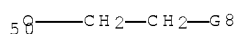
MSTR 1



G3 = 16



G4 = 50



G8 = NH2

Patent location: claim 1

Note: additional ring formation also claimed  
 Note: substitution is restricted  
 Note: also incorporates claim 13

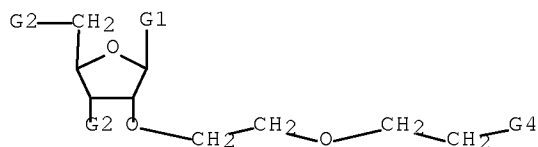
REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 24 OF 27 MARPAT COPYRIGHT 2008 ACS on STN  
 ACCESSION NUMBER: 132:176562 MARPAT Full-text  
 TITLE: 2'-O-Aminoethyloxyethyl-modified oligonucleotides and their use for inhibiting gene expression  
 INVENTOR(S): Manoharan, Muthiah; Cook, Phillip Dan  
 PATENT ASSIGNEE(S): Isis Pharmaceuticals, Inc., USA  
 SOURCE: PCT Int. Appl., 60 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 3  
 PATENT INFORMATION:

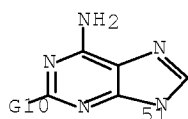
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000008044	A1	20000217	WO 1999-US17895	19990806
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US 6043352	A	20000328	US 1998-130566	19980807
CA 2339178	A1	20000217	CA 1999-2339178	19990806
AU 9954693	A	20000228	AU 1999-54693	19990806
AU 750469	B2	20020718		
EP 1102787	A1	20010530	EP 1999-940938	19990806
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
JP 2002522449	T	20020723	JP 2000-563677	19990806
PRIORITY APPLN. INFO.:			US 1998-130566	19980807
			WO 1999-US17895	19990806

AB 2'-O-Modified ribosyl nucleosides and modified oligomeric compds. containing such nucleosidic monomers are disclosed. Oligomeric compds. are disclosed that have increased binding affinity as shown by mol. modeling expts. The 2'-O-modified nucleosides of the invention include ring structures that position the sugar moiety of the nucleosides preferentially in 3' endo geometries.

MSTR 1



G1 = 51



G4 = 20



Patent location: claim 1

REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 25 OF 27 MARPAT COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 129:270610 MARPAT Full-text

TITLE: Antisense inhibition and diagnostic agents of angiogenin expression and inhibiting tumors associated with angiogenesis

INVENTOR(S): Fett, James W.; Olson, Karen A.

PATENT ASSIGNEE(S): President and Fellows of Harvard College, USA

SOURCE: PCT Int. Appl., 72 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

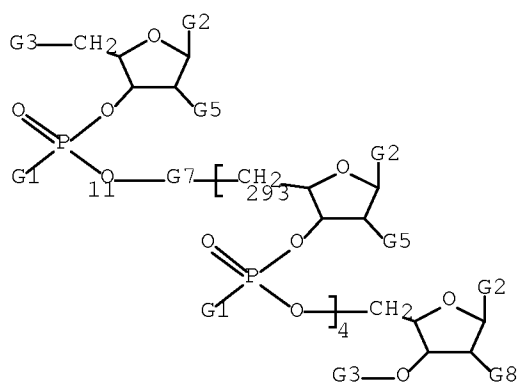
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PATENT INFORMATION:

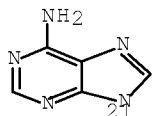
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9842722	A1	19981001	WO 1998-US5651	19980320
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RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
AU 9865782	A	19981020	AU 1998-65782	19980320
EP 970102	A1	20000112	EP 1998-911943	19980320
R: CH, DE, FR, GB, IT, LI				
US 6265388	B1	20010724	US 1998-45301	19980320
CA 2293591	A1	19981001	CA 1998-2293591	19980328
US 20020019359	A1	20020214	US 2001-863777	20010523
PRIORITY APPLN. INFO.:				
				US 1997-41182P
				US 1998-45301
				WO 1998-US5651

AB Disclosed are oligonucleotide compds. that inhibit the expression of angiogenin when administered to a mammal. Sense and antisense phosphorothioate oligodeoxynucleotides were designed based on the nucleic acid sequences of the angiogenin gene encompassing the AUG initiation codon and transcriptional start site regions, the 3'-termination site, and the 5'-TATA box site. Thus, phosphorothioated 5'-GCCCATCACCATCTCTTC-3' (JF2S) in combination with lipofectin inhibited in vitro production of angiogenin by 30-38% in PC-3 and HT-29 tumor cell types in comparison to treatment with lipofectin alone or with the control sense oligonucleotide. High doses of antisense JF2S also protected 50% of mice from forming regional lymph node metastasis in an orthotopic model of human prostate cancer metastasis in athymic mice. Also disclosed are methods and pharmaceutical compns. for inhibiting the expression of angiogenin useful in therapy or diagnosis.

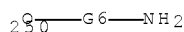
MSTR 1



G2 = 21



G5 = 250



G6 = (1-10) CH2

Patent location:

claim 13

REFERENCE COUNT:

6

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 26 OF 27 MARPAT COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 128:89088 MARPAT [Full-text](#)



TITLE: Preparation of 2'-substituted antisense oligoribonucleotide duplexes and triplexes as antitumors

INVENTOR(S): Cuenoud, Bernard; Altmann, Karl-Heinz; Martin, Pierre; Moser, Heinz

PATENT ASSIGNEE(S): Novartis A.-G., Switz.; Cuenoud, Bernard; Altmann, Karl-Heinz; Martin, Pierre; Moser, Heinz

SOURCE: PCT Int. Appl., 141 pp.  
CODEN: PIXXD2

DOCUMENT TYPE: Patent

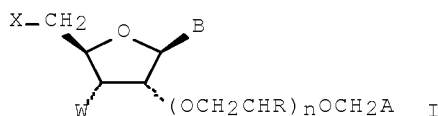
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9746569	A2	19971211	WO 1997-EP2738	19970527
WO 9746569	A3	19980219		
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
CA 2256765	A1	19971211	CA 1997-2256765	19970527
AU 9730297	A	19980105	AU 1997-30297	19970527
EP 906329	A2	19990407	EP 1997-924997	19970527
EP 906329	B1	20031008		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
JP 2000512630	T	20000926	JP 1998-500187	19970527
AT 251638	T	20031015	AT 1997-924997	19970527
ZA 9704952	A	19971208	ZA 1997-4952	19970605
US 6670468	B1	20031230	US 2001-753943	20010103
US 20050159374	A1	20050721	US 2003-696488	20031029
PRIORITY APPLN. INFO.:				
			CH 1996-1432	19960606
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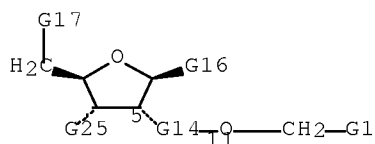
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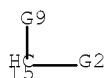
AB Antisense oligodeoxyribonucleotide duplexes and triplexes which comprises at least one nucleoside building block of the formula I (R = H, Me, Et, alkoxymethyl; n = 0-2; A = alkylamine, N-containing heterocycle; B = nucleobase; X, W = independently OH, internucleosidic bridge) were prepared as antitumors. Antitumor activities of antisense oligodeoxyribonucleotides were 9-42 expressed as T/C % (mean increase of tumor vols. of treated animals

divided by the mean increase of tumor vols. control animals multiplied by 100).

MSTR 1

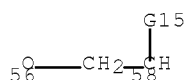


G1 = 15

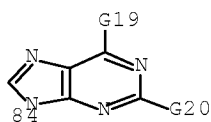


G2 = NH2

G14 = (0-2) 56-5 58-11



G16 = 84



Derivative: and salts  
Patent location: claim 1

L13 ANSWER 27 OF 27 MARPAT COPYRIGHT 2008 ACS on STN  
ACCESSION NUMBER: 124:261619 MARPAT Full-text  
TITLE: Solution phase synthesis of oligonucleotides.  
INVENTOR(S): Ravikumar, Vasulinga; Cole, Douglas L.  
PATENT ASSIGNEE(S): Isis Pharmaceuticals, Inc., USA  
SOURCE: PCT Int. Appl., 55 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 4

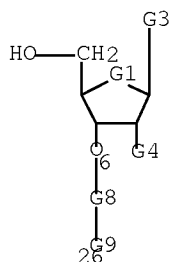
## PATENT INFORMATION:

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WO 9532980	A1	19951207	WO 1995-US6825	19950526
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RW:	KE, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG			
US 5571902	A	19961105	US 1994-249442	19940526
AU 9526570	A	19951221	AU 1995-26570	19950526
EP 766688	A1	19970409	EP 1995-921510	19950526
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE			
US 6294664	B1	20010925	US 1997-737875	19970117
PRIORITY APPLN. INFO.:			US 1994-249442	19940526
			US 1993-99075	19930729
			WO 1995-US6825	19950526
OTHER SOURCE(S):		CASREACT 124:261619		
GI				

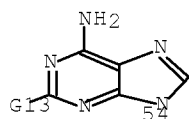
\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

AB Title compds. (I; Q = O, s, CH<sub>2</sub>, CHF, CF<sub>2</sub>; B = nucleosidic base; X = OH, SH, SMe, F, OCN, aminoalkoxy, alkoxy, alkyl, alkylaryl, aralkyl, Cl, Br, CN, CF<sub>3</sub>, OCF<sub>3</sub>, SMe, amino, polyalkylamino, aminoalkylamino, N<sub>3</sub>, substituted silyl, RNA cleaving group, conjugate, reporter group, intercalator, group for improving the pharmacokinetic or pharmacodynamic properties of an oligonucleotide, etc.; W, Y = protecting group; Z = O, S; T = phosphorus blocking group; n = 0-50), were prepared by reaction of synthon (II) with synthon (III; U = phosphite activating group). Thus, 5'-O-(4,4'-dimethoxytrityl)-3'-O-[(N,N-diisopropylamino)-2-(diphenylmethylsilyl)ethoxyphosphino]deoxyguanosine and tetrazole were stirred 15 min. in MeCN; 3'-(O-levulinyl)thymidine phosphorothioate dimer (preparation given) in MeCN was added. After 15 min the solution was cooled to 0° and treated with 3H-1,2-benzodithiol-3-one 1,1-dioxide to give 5'-(O-4,4'-dimethoxytrityl)-3'-(O-levulinyl)-dG-T-T phosphorothioate trimer.

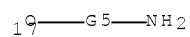
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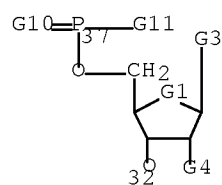
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G3 = 54



G4 = 17



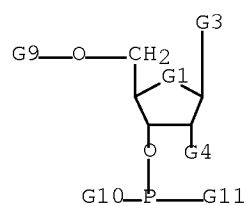
G5 = (1-10) CH2  
G8 = (0-5) 37-6 32-26



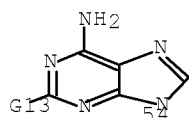
Patent location:  
Note:

claim 1  
G8 may contain a maximum of 50 nucleotide units

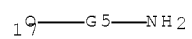
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G3 = 54



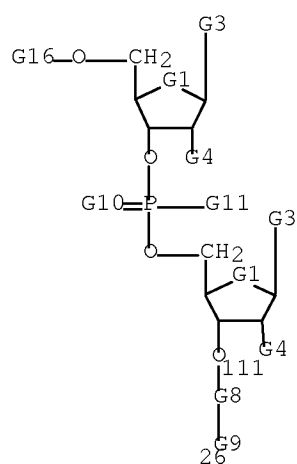
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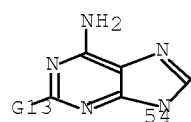
G5 = (1-10) CH<sub>2</sub>

Patent location: claim 1

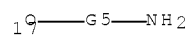
MSTR 3



G1 = O  
G3 = 54

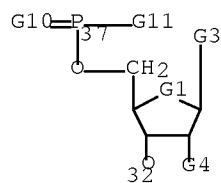


G4 = 17



G5 = (1-10) CH<sub>2</sub>

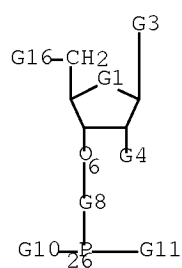
G8 = (0-4) 37-111 32-26



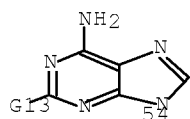
Patent location:  
Note:

claim 1  
G8 may contain a maximum of 50 nucleotide units

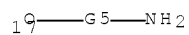
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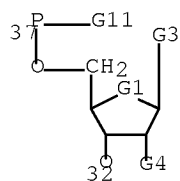
G1 = 0  
G3 = 54



G4 = 17



G5 = (1-10) CH2  
G8 = (1-5) 37-6 32-26



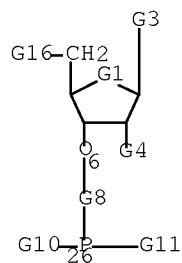
Patent location:

claim 14

Note:

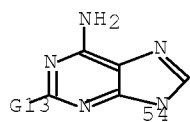
G8 may contain a maximum of 200 nucleotide units

MSTR 6

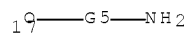


G1 = 0

G3 = 54

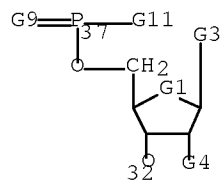


G4 = 17



G5 = (1-10) CH2

G8 = (1-5) 37-6 32-26



Patent location:

claim 14

Note:

G8 may contain a maximum of 200 nucleotide units

=> fil beilst

FILE 'BEILSTEIN' ENTERED AT 16:55:01 ON 23 MAY 2008

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FILE LAST UPDATED ON April 1, 2008

FILE COVERS 1771 TO 2008.

\*\*\* FILE CONTAINS 10.322,808 SUBSTANCES \*\*\*

>>>PLEASE NOTE: Reaction Data and substance data are stored in  
separate documents and can not be searched together in one query.  
Reaction data for BEILSTEIN compounds may be displayed  
immediately with the display codes PRE (preparations) and REA  
(reactions). A substance answer set retrieved after the search  
for a chemical name, a compounds with available reaction  
information by combining with PRE/FA, REA/FA or more generally  
with RX/FA. The BEILSTEIN Registry Number (BRN) is the link  
between a BEILSTEIN compound and belonging reactions. For mo  
detailed reaction searches BRNs can be searched as reaction  
partner BRNs Reactant BRN (RX.RBRN) or Product BRN (RX.PBRN).<<<

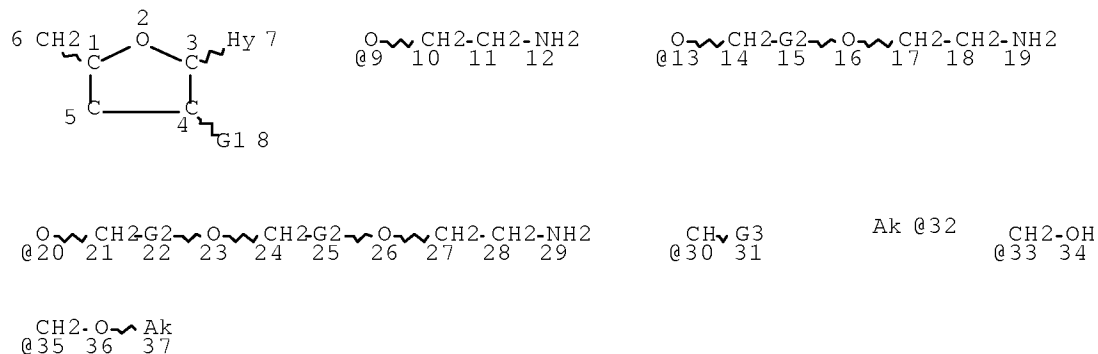
>>> FOR SEARCHING PREPARATIONS SEE HELP PRE <<<

\*\*\*\*\*  
\* PLEASE NOTE THAT THERE ARE NO FORMATS FREE OF COST. \*  
\* SET NOTICE FEATURE: THE COST ESTIMATES CALCULATED FOR SET NOTICE \*  
\* ARE BASED ON THE HIGHEST PRICE CATEGORY. THEREFORE; THESE \*  
\* ESTIMATES MAY NOT REFLECT THE ACTUAL COSTS. \*  
\* FOR PRICE INFORMATION SEE HELP COST \*  
\*\*\*\*\*

>>> Price change as of January 1st, 2008: Connect Time and Structure  
Search fees re-introduced. See NEWS and HELP COST <<<

=> d que 19

L1 STR



VAR G1=9/13/20

VAR G2=CH2/30

VAR G3=32/33/35

NODE ATTRIBUTES:

CONNECT IS E1 RC AT 32



CONNECT IS E1 RC AT 37  
 DEFAULT MLEVEL IS ATOM  
 GGCAT IS LOC SAT AT 32  
 GGCAT IS LOC SAT AT 37  
 DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:  
 RING(S) ARE ISOLATED OR EMBEDDED  
 NUMBER OF NODES IS 37

STEREO ATTRIBUTES: NONE  
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 L15 0 L9 AND 6509342/BABSAN

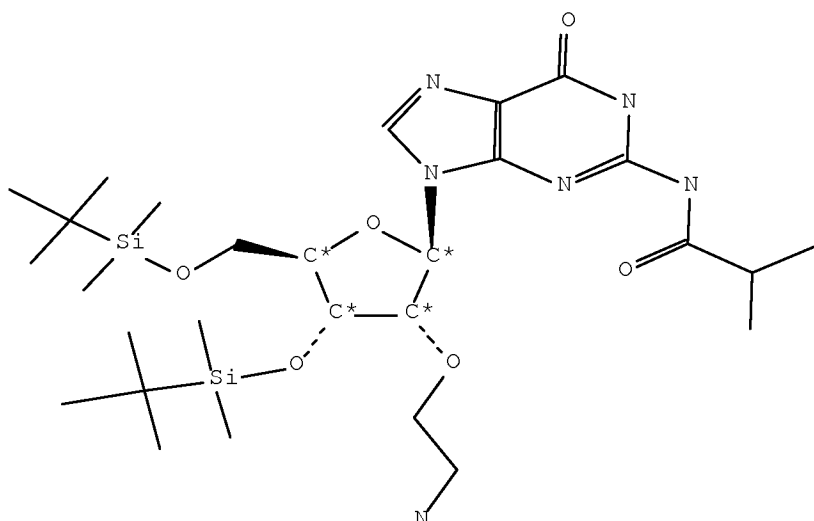
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 L17 3 L9 OR L14

=> d l17 ide allref tot

L17 ANSWER 1 OF 3 BEILSTEIN COPYRIGHT 2008 BEILSTEIN MDL on STN

Beilstein Records (BRN):	10135874
Chemical Name (CN):	N-<9-<3-(2-amino-ethoxy)-4-(tert-butyl-dimethyl-silanyloxy)-5-(tert-butyl-dimethyl-silanyloxymethyl)-tetrahydro-furan-2-yl>-6-oxo-6,9-dihydro-1H-purin-2-yl>-isobutyramide
Autonom Name (AUN):	N-<9-<3-(2-amino-ethoxy)-4-(tert-butyl-dimethyl-silanyloxy)-5-(tert-butyl-dimethyl-silanyloxymethyl)-tetrahydro-furan-2-yl>-6-oxo-6,9-dihydro-1H-purin-2-yl>-isobutyramide
Molec. Formula (MF):	C28 H52 N6 O6 Si2
Molecular Weight (MW):	624.93
Lawson Number (LN):	30733, 20554, 3798, 3777, 3122, 1174
File Segment (FS):	Stereo compound
Compound Type (CTYPE):	heterocyclic
Constitution ID (CONSID):	8519549
Tautomer ID (TAUTID):	9480720
Entry Date (DED):	2006/02/02
Update Date (DUPD):	2006/02/02



## Field Availability:

Code	Name	Occurrence
BRN	Beilstein Records	1
CN	Chemical Name	1
AUN	Autonomname	1
MF	Molecular Formula	1
FW	Formular Weight	1
LN	Lawson Number	6
FS	File Segment	1
CTYPE	Compound Type	1
CONSID	Constitution ID	1
TAUTID	Tautomer ID	1
ED	Entry Date	1
UPD	Update Date	1

## This substance also occurs in Reaction Documents:

Code	Name	Occurrence
RX	Reaction Documents	2
RXREA	Substance is Reaction Reactant	1
RXPRO	Substance is Reaction Product	1

## All References:

ALLREF

- Jin, Shengxi; Miduturu, Chandrasekhar V.; McKinney, David C.; Silverman, Scott K., J. Org. Chem., CODEN: JOCEAH, SIR70(11), <2005>, 4284 - 4299; BABS-6509342

L17 ANSWER 2 OF 3 BEILSTEIN COPYRIGHT 2008 BEILSTEIN MDL on STN

Beilstein Records (BRN): 10133679  
 Chemical Name (CN): N-<9-<3-(2-amino-ethoxy)-4-(tert-butyl-

dimethyl-silanyloxy)-5-(tert-butyl-  
 dimethyl-silanyloxymethyl)-tetrahydro-  
 furan-2-yl>-9H-purin-6-yl>-isobutyramide  
 Autonom Name (AUN): N-<9-<3-(2-amino-ethoxy)-4-(tert-butyl-  
 dimethyl-silanyloxy)-5-(tert-butyl-  
 dimethyl-silanyloxymethyl)-tetrahydro-  
 furan-2-yl>-9H-purin-6-yl>-isobutyramide  
 Molec. Formula (MF): C28 H52 N6 O5 Si2  
 Molecular Weight (MW): 608.93  
 Lawson Number (LN): 30692, 20554, 3798, 3777, 3122, 1174  
 File Segment (FS): Stereo compound  
 Compound Type (CTYPE): heterocyclic  
 Constitution ID (CONSID): 8517443  
 Tautomer ID (TAUTID): 9479728  
 Entry Date (DED): 2006/02/02  
 Update Date (DUPD): 2006/02/02

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## Field Availability:

Code	Name	Occurrence
BRN	Beilstein Records	1
CN	Chemical Name	1
AUN	Autonomname	1
MF	Molecular Formula	1
FW	Formular Weight	1
LN	Lawson Number	6
FS	File Segment	1
CTYPE	Compound Type	1
CONSID	Constitution ID	1
TAUTID	Tautomer ID	1
ED	Entry Date	1
UPD	Update Date	1

This substance also occurs in Reaction Documents:

Code	Name	Occurrence
RX	Reaction Documents	2
RXREA	Substance is Reaction Reactant	1
RXPRO	Substance is Reaction Product	1

## All References:

## ALLREF

- Jin, Shengxi; Miduturu, Chandrasekhar V.; McKinney, David C.; Silverman, Scott K., J. Org. Chem., CODEN: JOCEAH, SIR70(11), <2005>, 4284 - 4299; BABS-6509342

Beilstein Records (BRN): 8373330  
 Chemical Name (CN): 2'-O-<2-(amino)ethyl>-5'-O-(4,4'-dimethoxytrityl)-5-methyluridine  
 Autonom Name (AUN): 1-<3-(2-amino-ethoxy)-5-<bis-(4-methoxy-phenyl)-phenyl-methoxymethyl>-4-hydroxy-tetrahydro-furan-2-yl>-5-methyl-1H-pyrimidine-2,4-dione  
 Molec. Formula (MF): C33 H37 N3 O8  
 Molecular Weight (MW): 603.67  
 Lawson Number (LN): 28796, 20554, 6582, 3122, 289  
 File Segment (FS): Stereo compound  
 Compound Type (CTYPE): heterocyclic  
 Constitution ID (CONSID): 7108715  
 Tautomer ID (TAUTID): 7887954  
 Entry Date (DED): 2000/03/08  
 Update Date (DUPD): 2008/01/25

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

Field Availability:

Code	Name	Occurrence
BRN	Beilstein Records	1
CN	Chemical Name	1
AUN	Autonomname	1
MF	Molecular Formula	1
FW	Formular Weight	1
LN	Lawson Number	5
FS	File Segment	1
CTYPE	Compound Type	1
CONSID	Constitution ID	1
TAUTID	Tautomer ID	1
ED	Entry Date	1
UPD	Update Date	1
NMR	Nuclear Magnetic Resonance	3

This substance also occurs in Reaction Documents:

Code	Name	Occurrence
RX	Reaction Documents	3
RXREA	Substance is Reaction Reactant	2
RXPRO	Substance is Reaction Product	1

All References:

ALLREF

1. Prakash, Thazha P.; Pueschl, Ask; Manoharan, Muthiah, Nucleosides, Nucleotides & Nucleic Acids, CODEN: NNNAFY, 26(2), <2007>, 149 - 159; BABS-6693986
2. Prakash, Thazha P.; Pueschl, Ask; Lesnik, Elena; Mohan, Venkatraman; Tereshko, Valentina; Egli, Martin; Manoharan, Muthiah, Org. Lett., CODEN: ORLEF7, 6(12), <2004>, 1971 - 1974; BABS-6459911
3. Manoharan, Muthiah; Prakash, Thazha P.; Barber-Peoc'h, Isabelle; Bhat,

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Balkrishen; Vasquez, Guillermo; Ross, Bruce S.; Cook, P. Dan,  
J.Org.Chem., CODEN: JOCEAH, 64(17), <1999>, 6468 - 6472; BABS-6182585  
4. Manoharan, Muthiah; Prakash, Thazha P.; Barber-Peoc'h, Isabelle; Bhat,  
Balkrishen; Vasquez, Gulliermo; Ross, Bruce S.; Cook, P. Dan,  
Nucleosides Nucleotides, CODEN: NUNUD5, 18(6-7), <1999>, 1199 - 1202;  
BABS-6181867

=> d his nofil

(FILE 'HOME' ENTERED AT 16:36:34 ON 23 MAY 2008)

FILE 'REGISTRY' ENTERED AT 16:36:58 ON 23 MAY 2008

L1 STR  
L2 0 SEA SSS SAM L1  
L3 4 SEA SSS FUL L1  
D SCA

FILE 'CAPLUS' ENTERED AT 16:43:55 ON 23 MAY 2008

L4 9 SEA ABB=ON PLU=ON L3

FILE 'WPIX' ENTERED AT 16:45:52 ON 23 MAY 2008

L5 0 SEA SSS SAM L1  
L6 1 SEA SSS FUL L1  
L7 1 SEA ABB=ON PLU=ON L6/DCR

FILE 'BEILSTEIN' ENTERED AT 16:46:14 ON 23 MAY 2008

L8 1 SEA SSS SAM L1  
L9 3 SEA SSS FUL L1  
SEL BABSAN L9

FILE 'BABS' ENTERED AT 16:47:16 ON 23 MAY 2008

L10 5 SEA ABB=ON PLU=ON (6509342/AN OR 6181867/AN OR 6182585/AN OR  
6459911/AN OR 6693986/AN)  
D 1-5

FILE 'MARPAT' ENTERED AT 16:48:01 ON 23 MAY 2008

L11 1 SEA SSS SAM L1  
L12 17 SEA SSS FUL L1

FILE 'CAPLUS' ENTERED AT 16:49:03 ON 23 MAY 2008

D QUE L4

FILE 'WPIX' ENTERED AT 16:49:13 ON 23 MAY 2008

D QUE L7

FILE 'MARPAT' ENTERED AT 16:49:19 ON 23 MAY 2008

D QUE L12

FILE 'CAPLUS, WPIX, MARPAT' ENTERED AT 16:49:24 ON 23 MAY 2008

L13 27 DUP REM L4 L7 L12 (0 DUPLICATES REMOVED)  
ANSWERS '1-9' FROM FILE CAPLUS  
ANSWER '10' FROM FILE WPIX  
ANSWERS '11-27' FROM FILE MARPAT  
D L13 IBIB ABS HITSTR 1-10  
D L13 IBIB ABS QHIT 11-27

FILE 'BEILSTEIN' ENTERED AT 16:55:01 ON 23 MAY 2008

D QUE L9

L14 2 SEA ABB=ON PLU=ON L9 NOT BABSAN/FA

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L15      0 SEA ABB=ON  PLU=ON  L9 AND 6509342/BABSAN
L16      0 SEA ABB=ON  PLU=ON  L9 AND 6181867/BABSAN
L17      3 SEA ABB=ON  PLU=ON  L9 OR L14
          D L17 IDE ALLREF TOT
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